	2328	Qy 2346 leArgLysAlaLeuMetGlyLysTyrAspG] :: :: Db 10246 TGACAAGGGCC	Qy 2366 snalaPheAsnProLeuAsnAlaSerAlaSe : Db 10282 CCACTTGGGAAGGACGGGATCCTCACTG	2386	10342	OY 2406 Serialyargeroserserarguysalalyss	Oy 2418 AlaProGlyLeuAlaSerGlyAspArgProl	Qy 2438 AspCysAsnArgArgThrProleuThrAsnJ Db 10453CCTCCACTC	Qy 2458 GlySerThrProPheProTyrAsnProLeul	10486	Qy 2478 SerProProProBroBycelyLeuProAlaGly6		DB 10558CCCGCACCCCTGCAGTGC	Search completed: April 16, 2004, 02:38:23 Job time : 14964 secs							
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          74692 GCGTCGGCTTCTATCCCGGATACGTGAGTGAACACAGCTCGACCGAGACCAACGCGGCAA
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                                                                                                                                                                                                                                                                                                    The invention relates to 446 newly isolated polynucleotide sequences. The activity of polynucleotides of the invention may be described as, vulnerary, neuroprotective, immunomodulator, cytostatic and anti-inflammatory. Compositions comprising nucleic acids of the invention are useful for treating a mammalian subject, or as nutritional sources or supplements. These are useful in gene therapy, particularly for treating wounds, burns or ulcers, Alzheimer, disease, Huntington's disease, amyotrophic lateral sclerosis, autoimmune disorders, cancer or inflammation. The nucleic acids and polypeptides are also useful in diagnostic and research methods. The sequences given in records ABQ60788-ABQ61233 represent polynuclectides of the invention. NOTE: The sequence data for this parent did not form part of the printed specification, but was obtained in electronic formet directly from WIPO at
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            -----calcicrradalcalcacacicacadas
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                                                                                                                                                                                  New polypeptides and their encoded proteins, useful as nutritional sources or supplements, or in gene therapy, particularly for treating wounds, Alzheimer's disease, amyotrophic lateral sclerosis, cancer or
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Best Local Similarity:
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Yang Y,
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qq	78590GCAAACCCCGAAACGCCGGCCCATACCCCTGCGCCCG- 78549	Ov 1490 uAs	1490 uAspValMetAla
δ	1145 alGlyProValThrMetGlyLeuProLeuProMetAspProLysLysLeuAlaProPheS 1165	7	
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ò	1165 erGlyValLysGlnGluGlnLeuSerProArgGlyGlnAlaGlyProProGluSerLeuG 1185	7	
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ò	1242 ThrileThrArgllelleGlyGluAspSerProSerArgLeuAspArgGlyArgGluAsp 1261	77552	TCCTGCGC
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ò	1262 SerLeuProLysGlyHisValIleTyrGluGlyLysLysGlyHisValLeuSerTyrGlu 1281	7	TCTCGCCG
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ò	1282 GlyGlyMetSerValThrGlnCysSerIysGluAspGlyArgSerSerSerGerGlyProPro 1301	7	
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ò	1302 HisGluThrAlaAla	77420	 ccaactic
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à	1318 ValGlyArgAlaIleSerSerAlaSerIleGluGlyLeuMetGlyArgAlaIleProPro 1337	77360	ACCGCACC
qq	78140 CCGAGTCGGGCCGTTGCACCACTGCAGCCAACAATCCATGCTCGGCGCCCCACA 78087		1680 uAsnArqGlnThı
à	1338 GluArgHisSerProHisHisLeuLysGluGlnHisHisIleArgGlySerlleThrGln 1357	7	1 1 1
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δ	1358 GlyIleProArgSerTyrValGluAlaGluGluAspTyrLeuArgArgGluAlaLysLeu 1377	77315	CGTCCCGAAGCG
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'n	1378 LeuLysArgGluGlyThrProProProProProSerArgAspLeuThrGluAla-Ty 1397	77270	
a a	78020 ACGCGTACGTCGGCAACTCCACCCGGCGGCCTCCCAGGTCGGCAAACGCGGCACGCCAGGT 77961		1210 0000000000000000000000000000000000
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ò	1433 uArgHisThrProGluLeuProLeuAlaProArgProLeuLysGluGlySerIleThrGl 1453	Db 77155 CCCGGGTGCGCC	 cggTgcG
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8Hi8AspValArgSerLeulleGlySerProGlyArgThrPhePr 	ProProvalHisProLe 1490
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GGTCTCGGCAACAACGTCCTGGCC 9726 LysLysLeuAsnThrHisAsnArgA 2300 9054 ACCAAGACAGCTGCTCAGGGCCCTC 9157 AlaAsnAlaPheAsnProteuAsnA 2373 GIGCAGICCACACAGCCTGCCCA-G 9216 AlaMetProlleThrAlaAlaAspG 2388 LeuGlnAla---- 2473 ccgdcccagcrcacacacrcagr 9097 lleArgLysAlaLeuMetGlyLysT 2354

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d Q	6592 CCACACCAGAGCCCCCCTACTAAGGTGACAGAGTGGATCACAAGGCAGGAGG 0042	λō	1851 oAlaSerHisAlaHisAlaHisGlnHisSerProlleSerProArgThrGlnAspAlaLe 1871
ð i	ProPheAlaGlyHisLeuProArgGlySerFrovalIII.mechigGlyHislard	qq	CAPACAGAGAGGGGGTGCTAATGAAACAGTCGGTTCCACCCAGGGTCCATGCCTGGAACAGAGTCGAACAGAGTCCTATGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTG
දු දි		Š t	1871 uGlnGlnArgbroSerValLeuHisAsninrd-lyweLuysety.reiler
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ò	1586 ThrSerThrProArgGluIleAlaLysSer-ProHisSerThrValProGluHisHisPr 1605	g qq	
qq		λŏ	1908 gProAlaAlaThrPheProProAlaThrHisCysProLeuGlyGlyThrLe 1925
ò	luHisLeuLeuArgGlyValSerGlyValAspLeuTyrAr	qq	
qa	0	δλ	1925 uAspGlyValTyrProThrLeuMetGluProValLeuLeuProLysGluAlaProArgVa 1945
ð.	gSerHisileBroLeuAlaPheAspProInfSetileFroAigG-yileFroLeus-Fra- 	q	GGACATTGAATTTCAGCAGTCAGTGTCCAAGTCCCAGGTCAAACCTGATTCTGTCACAGC
음 8	ArgHi	ð í	
3 8	: CCTTTAGAAGAAAAAACAGCACCTCCAGTGACA	g &	1962
ò		3 20	TTCCACGTTGGTACTGACCGCCCAGACATATAA
Ор		λō	1970 uProAlaSerSerProSerLysGlySerGluProArgProLeu 1984
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ò		q	ACCCAGGGGATCAACACACCCCCTGT 794
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ò	621 GluThrAla 623	q	4222 GTAGAC
ପୁ	3160 CAGGATGCTGGCAGATITGATGTGAGTTTCCCAAACAGCATAATTAAGAGAGATAGCCTT 3219	λō	761 AspThr
ò	623 623	q ₀	::: 4282 GAGGAA
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626 (GlyLeuLeuGluHisGlyArg
638	AlaArgMetValGlySerLysThrValSerGln 648
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3700	TTTGCTTCTCGTTTTTTACACAGCTCAATCTTTGAACAAGATTCCAAGCGATTGCAGCAT 3759
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3820	TCTGAGGGAGGAAACAGCACAACTGATTCCATTCAAGAACCAGTAGTTCTGTTCCATAGC 3879
671	685
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3940	GAGGTTGAGAAACAGGAAGATACAGAGAATCATCCCAAAACCCCCAGAATCTGCTCCTGAG 3999
689	700
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701	707 1 1 1 1 1 1 1 1 1
4060	ACTCTAGAATCAGCCCCATCAGCACTAGAGAACACCACTGGTĠACAAAACGGTAGAGGCG 41
708	SerGlyValSerGlyAsnGluGluGluMetValGluGluAlaGluAlaLeuHisAlaSer 727
4120	CCTTTGGTAACAGAAGAGAACTGTGGAGCCAGCTACCGTCTCA 41.
728	GlyAsnGluValProArgGlyGluCysSerGlyProAlaThrValAsnAsnSerSerAsp 747
4165	GAAGAAGCAAAGCTGCATCTGAACCTGCTCCTGCCCTGTGGAACAGCTGGAACAA 42
748	ThrGluSerIleProSerProHisThrGluAlaAlaLy8760
4224	
761	AspThrclyGlndsnGlyProUySProProAlaThrLeUclyAldaspc1yFlorLorlor
781	GlyProProThrProProArgArgThrSerArgAlaPro1leGluProThrPro 798
4327	CCTCCAACTCCCGGGGCCTCGTTTTCCCAGGCAGGAAGGA
799	AlaSerGluAlaThrGlyAlaProThrProProProAlaProProSerProSerAlaPro

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                                                                                        291 GAACGAAGTCAAAGTCCTGTTCACTTGCGACGTCCACAGAGTCCTGGAGCGTCTCCCTCT 1350
                                                                                                                                                                                1351 CAGGCAGAGAGTTGCCGAGTGATTCTGAGAGGAGGCTTTACAGCCGATCCTCAGACCGG 1410
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| S25 AGAGTGAGAGAGAGAGCGCTTAATACGGAAGGAAAAAGTGGAAAAAGGACAAAAACTGAC 1584
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|1885 CAGACGGAGCCTGCAAAATCTGATTGTCTAAACTGGAATCAGTTAGAATGAAAGTACCA 1944
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                                                                                                                                                                                                                                                                                                                                                                                                               ArgleuSerLysGluGluLeuIleGlnAsnMetAspArgValAspArgGluIleThrMet 185
                                                                                                                                                                                                                                                           |||| AGT-----GGAAGCTGTAGCTCCTCCCCTCCAAGATATGAGAAACTGGACAAGTCT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         262 AspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLysLysLeu---
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                                                                                                                                   --AspLeuThrLysAspArg
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                                             GluLeuLeuProAspPro-----LeuLeuArgPro-
                                                                                                                                     124 LeuAlaThrGlyGlnProAlaGlySerGlu-
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     GACTATAACCAAGATCGTACATATTATGAGAGTGTTCGAACTCCAGGCACTTATCCTGAG 1056
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         cartilage damage, nocturnal haemoglobinuria, antiinflammatory disease; to enhance coagulation; to inhibit thrombosis; and as a contraceptive
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   -----Leu 110
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AAC74446 to AAC77606 encode the proteins given in AAB40237 to AAB43397, which represent the human ORFX open reading frames 1 to 3161. The ORFX sequences have activities such as: cytostatic; hepatotropic; vulnerary;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     61 ProGlnArgArg---ArgProSerLeuLeuSerGluPheGlnProGlyAsnGluArgSer
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                                                                                                                                                                                                                                                                                                                                                               Novel nucleic acids and peptides derived from open reading frame X, useful for treating e.g. cancers, proliferative disorders, neurodegenerative disorders and cardiovascular disease.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 11680 BP; 3290 A; 3113 C; 2858 G; 2417 T; 0 U; 2 Other;
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368
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Conservative:
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99US-0127636P.
99US-0127728P.
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                                                                                                                                     05-APR-1999;
30-MAR-2000;
                                             31-MAR-2000;
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02-APR-1999;
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Pred. No.:
05-OCT-2000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1117
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DB:
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		Qy 2330
ò	2092 lyProVal	DP 65968 ACAGATTCCCGATTTCG
d d	68978 GTCCCATCGGCGCACCGCATAGGCCTTACAACGACCATCGGCGGACAG 66929	2341
ò	enb	65908
qa	66928	2361
ò	2112 erGlnProSerSerSerProLeuGlnThrAlaPro	
Ор		Qy 2378 roAlaAlaMetProlle
ò		Db 65794 CCAGATACTGCTCGACC
q		Qy 2398 ly
ò	2135 eualaGinHisile	Db 65737 CCGAGCCGACGTCGTAI
q	CCCAACACATACGCCACCCCGGAGGCCACGCTCAGGCTCGCCTCGAGGCCAGGCCCC	
<i>&</i> :		
දු දි	GCTCCAGGTCACCCGGGCCTTGGGCCCCAAAAAAAAAAA	
<u> </u>	AACACGCCGTCTGCGAACCCGCAACGTGATTGGGTCGA	65617
ŀò	2165 erPhePro 2167	ATCCCGGATACGT
අධ	 66582 GCTTCCCACACACTTCCAACAGCAACCGCTGCTGGGGATCCATCGCAAGCGCCTCGCTG 66523	2447
à	2168GlyAlaSerCysProValLeuAspLeuArgArgProProSer 2181	65497
qa	GAACCCGCCGCAC	Qy 2467 LeuIleMetArgLeuG
ò	31yA1	Db 65440 GCCCGAACAGCGGCAC
đ		Qy 2486 a
ò	2192 aProAlaArgGlySerProHisSerGluGlyGlyLysArgSerProGluPr 2209	Db 65380 CATCCAGCGACGGCAC
qq		RESULT 98
ò	2228	AAC76700 ID AAC76700 standard; cDNA;
q	66348 CACAGGGTCTCCGGCGAATCCACCCCACCGGATAGCGGCATCCCATCCCCACCACCGCC 66289	XX AC AAC76700;
ò	alTyrProLe	XX DT 08-FBB-2001 (first entry
qq	!	XX DE Human ORFX ORF2255 polynu
ò	2244 uLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlySerLysSerProGly 2263	
qq	-	KW anticottiniant; cateopath
ò		
q		
ò	2280 ralametValLysSerLysLysLys-GlnGlulleAsnLysLysLeuAsnlhrHisAsnA 2299	
Q D		severe combi
ò	2299 rgAsnGluProGluTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetFroA 2318	KW thrombosis; contraceptive
a :	66067 AALAGAAGCCIGGGCCCAIGGCGAICAICAACGACGACAACAACAACAACAAAAAAAA	Homo sapiens
중 A		XX PN WO200058473-A2. XX
!		

frame, ORFX; detection; cytostatic; hepatotropic; atic; antiparkinsonian; nootropic; neuroprotective; copathic; antiarthritic; immunosuppressant; cardiant; ombolytic; coaquiant; exactropic; antidiabetic; ombolytic; coaquiant; vesteropic; antidiabetic; logical; immunosuppressive; antiinflammatory; rial; antifungal; antirheumatic; antithyroid; erapy; cancer; proliferative disorder; hypertension; esorder; osteoarthritis; graft vs host disease; se, diabetes mellitus; hypothyroidism; SCID; AIDS; orage; systemic lupus erythematosus; infection; nodeficiency; malaria; autoimmune disorder; asthma; aemia; nocturnal haemoglobinuria; burn; wound; ge damage; antiinflammatory disease; coagulation; ptive; SS. ||| ::::||| | ||| CGTCGGCGCGCTTCACACCTGCGGCCACCA 65618 :: CAGCAAACAACGCACTGCCGACAAGGCCCGGCAGCGTCGGCTTGT 65558 || ||||||| ::: |||| GTGAACACAGCTCGACCAACGCGGCAACGCCGTCCGGGG 65498 CCCTCGGACATCGGCGCGCAGCCCCAGCCGCGGCGATCCGGG 65381 ATCCCATGGTGCGCAGCGAGTCGGGGTACAGATGGCTGCGCGAAT 65678 GCCCAACGAGGGCCACGAGAGCTGGGGGGACAGGT 65909 : :::::||| GGCGTTGATGATCTTGGGTCCC---CCGGGATGGGTGACCCAGG 65852 -grecreageccerecgaaeccaageagergergacgicairec 65795 AspArgProSerSerAlaGlySerThrProPheProTyrAsnPro 2466 GlnAlaGly-ValMetAlaSerProProProGlyLeuProAl 2486 :::||| ::: ||| ||| |||| erhraljajjaAspGlyargSeraspHisThrLeuThrSerProG 2398 lyLysaljaLysValSerGly-----ArgproSerSerArgL 2413 laProGlyLeuAlaSerGlyAspArgProProServalSer---- 2431 eArgLysAlaLeuMetGlyLysTyrAspGlnTrpGluGluSerP 2361 ------GlnAlaValGlnGluHisAlaSerThrAsnMetG 2341 erValHisSerGluGlyAspCysAspArgArgThrProLeu-Thr nAlaPheAsnProLeuAsnAlaSer-----AlaSerLeuP cceccagiccediaaccereecediaarcaacac 65332 ----GlySerGlyProLeuAlaGlyProHisHis 2496 nucleotide sequence SEQ ID NO:4509. 11680 BP 2330 ò

Page 390

9 & 9 10 & 10	
6 8 6	1165 erGlyvalLysGlnGluc
g &	 69338 GCCAACCCCAACACGC 1203 alProGly-GlySer
7 d	69287 CGCCTGGCCGGCAGCCG
දු පු	1221 rAlaileThrTyrArgG :: 69227 CACCACCACCTGGACCC
ò	
qq	
දු දු	1261 pSerLeuProLy8GlyH 69193
8 8	
3 8	69185 CGTGCCTGCAACGTCA
ģ	1294
QQ	69125 GCCAACTCAACAAACC
ò	1305 aAla
음 2	69065 AGCCACGCCGCAGCGAC
; 음	
ò	1341 rProHisHisLeuLys
а	68951 CCCAGATCGCCGC
ζ	· 1361 gSerTyrValGluAla(
q	68902ĠĊĠ
ò	1381 uGlyThrProProPro
ф	68885 GGCAACTCCACCCGG
à i	1401 la
e è	1417 hrvallvegluglagl
5 음	68765 CCCAGCGT
ò	1437 roGluLeuProLeuAl
q	68736
ò	1457 euLysTyrAspThrGl
	6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8

AA199683 11 1100001 1 AA199683 12 1200001 1 AA199683 14 1400001 1 AA199683 15 1500001 1 AA199683 15 1500001 1 AA199683 19 100001 1 AA199683 20 100001 1 AA199683 30 100001 1 AA19	9683_40	US-09-522-753-5 (1-2517) x AA199683_18 (1-110000) QY	Db 72182 TTTGTGAGCGTTGATATCGACACTGCTGGAACGACCCCACACCCCGCGCATTGCTGAT 72123
cc are useful in diagnostics as expressed sequence tags (EST) for identifying expressed genes or for physical mapping of the human genome, cn in identifying mutations cresponsible for genetic disorders and other traits. The nucleotide responsible for genetic disorders and other traits. The nucleotide sequences are also useful as hybridisation probles, as oligomers for PCR, cording one mapping, in the recombinant production of creating antibodies specific for it. The present sequence is useful for generating antibodies specific for it. The present sequence creating antibodies specific for it. The present sequence creating isolated CDNA, BCT sequences. Note: The present sequence data contained in electronic format directly from USPTO at containing in electronic format directly from USPTO at containing in electronic format directly from USPTO at sequence 499 BP; 183 A; 80 C; 123 G; 111 T; 0 U; 2 Other; Alignment Score: Alignment Scores: Alignment Scores: Alignment Scores: Best Local Similarity: Alignment Scores: Best Local Similarity: Alignment Scores: Best Local Similarity: Best Local S	00 AlaarglysGlnTrpLysGlnLysPheCysGlnArgTyrAspGlnLeuwetGlua	G—re G—r G—r G—r 33 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3 3	Oy 410 earth

	RESULT 96 ACH32484 standard; CDNA; 499 BP. XX AC ACH32484; XX DT 13-OCT-2003 (first entry) XX DY 13-OCT-2003 (first entry) XX DY Whman endothelial cell cDNA #617. XX	XX XX XX PN 17-APR-2003. XX XY PF 30-JUL-2001; 2001US-00918995. XX XX PN 30-JUL-2001; 2001US-00918995. XX PN 4 (LABA/) LABATI PA (STAC/) STACHE-CRAIN B. PA (GTAC/) STACHE-CRAIN B. PA (JONE/) JONES L W. XX	Drmanac RT, Labat I, Stache-Crain B, Dickson MC, Jones LW; XX XX WPI; 2003-615964/58. XX PT as hybridization probes, as oligomers for PCR, for chromosome and gene PT as phybridization probes, as oligomers for PCR, for chromosome and gene PT mapping, in the recombinant production of protein, or in generating PT antisense DNA or RNA. XX PX Claim 1; SEQ ID NO 19696; 44pp; English. XX CC 38043 acDNA sequences, appearing as ACH12789-ACH50811, whose sequence was determined by the technique of SBM (sequencing by hybridisation). Also CC included is a purified polypeptide comprising a sequence corresponding to a reading frame of the novel polynucleotide. The nucleic acid sequences
InThralaProGlyValLysGlyHisGlnArgValValThrLeuAlaGlnHisIleSerG 2141	2224 roValSerProProGluGlyMetThrGluProGlyHisSerArgSerAlaVal 2241 9902 CTGCCAAGACACCAAGCCCCACCCCCACCCCTGCCCCGTCC 9961 2242TyrProLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgM 2257 9962 CTGTCCCCTTCCTGCCCTTGCTCCTCTGCCCCTAGTGAGTG	10063	2346 lehrgLyshlaLeuMetGlyLysTyrAspGlnTrpGluGluSerProProLeuSerh 2365 5 ::

	1863 eSerProArgT	8559 CCACCCAGGGT	1883 tLysGlyIleI	 8608 -TCAGGGGGG	1903ThrS	GCAGAA	1917 rHisCysProL	8727 T	1937 uLeuProLysG	::: 8769 GGTCAAACCTG	1955 rGlyHisAlaP	8829 AGGCTATGCGA	1962 sProproAlaA	8889 CTCTCTGTGA	1982 gProLeu	::: 8943 GCCCATTCACC	1985	9003 CCAGGGGATCA	1999	9063 TGTCACCACA	2000	9123 TCAGCCGGCC	2013 aProProAlaS	::: 9171 TCTGCCCAGC	2031 eSer]	9231 AGATCGAACTC	2049 rPro-GluGl)	- :: 9279 TCCTCGACCA	2065 hrHisAspLy	9339 TGCAT	2085 luLeuArgPro	9344CTACGGCG	2105 isLeuArgPr	9402 GTTCATCTCC	2119 euLeu	9462 GACTGTGTCC
qa	ò	q	δy	qq	ð	: A	ò	· 유	ò	셤	ò	q	8	qa	ò	qq	ò	: A	ò	; <u>a</u>	ò	셤	ò	: A	8	: 名	ò	අය	ò	qu	ò	qq .	ò	qa	ò	qu
		1 InrMetArgGluProInrFroArgLeuGinGluGiyseLieuseLseLseL 	7732 GCTCAGICTACTCCCAGCTCTTCCCCCAGACACACAAAAGCCTCTGAIGTTGACACC 7791	1578 LysalaSerGlnAspArgLysLeuThrSerThrProArgGluIleAlaLysSer-ProHi 1597	7792 AGCTCCAGCACCCTGAGGAAGATTCTCATGGACCCCAAGTATGTGTCTGCCACA 7845	1597 sSerThrValProGluHisHisProHisProIleSerProTyrGluHisLeuLeuArgGl 1617	7846 AGTGTCACTTCCACAAGTGTCACCACAGCCATTGCAGAGCCTG7888	1617 yValSerGlyValAspLeuTyrArgSerHisIleProLeuAlaPheAspProThrSerIl 1637			7921 CCCCGGCAGTTGACTCTAAAAGCCTTTAGAAGAAAAAAGGCACCTCCAGTGACAAAC 7980		aactctgagatacaagcetcggaggtgctggtagctgctgacaaggaaaaggtggctcca		8041 GTCATTGCTCCCAAAATTACCTCTTATTAGCCGGATG-CCTGTCAGCATTGACCTGGA 8099	uAsnArgGlnThrIleIleAsnAspTyrIleThrSerGlnGlnMetHisHisAsnThrAl 	8100 AAATTCACAGAAGATA 8115	1700 aThralaMetAlaGlnArgAlaAspWetLeuArgGlyLeuSerProArgGluSerSerLe 1720	8116 -ACCTIGGCAAAACCAGCICCICAAACCCICACIGGICIG	1720 uAlaLeuAsnTyrAlaAlaGlyProArgGlyllelleAspLeuSerGlnValProHisLe 1740	8155GIGAGCGCACTCACTGGCCTGGTCACGTCTCCCTGGTCCCGGT 8198	1740 uProValLeuValProProThrProGlyThrProAlaThrAlaMetAspArgLeuAlaTy 1760	8199 GAATGCCCTGAAAGGCCCCGTGAAGGCTCAGTGACCACACTGAAAGTTTGGTGAGC 8256	1760 rLeuProThrAlaProGlnProPheSerSerArgHisSerSerSerProLeuSerProGl 1780	8257ACCCTIGC 8264	1780 yGlyProThrHisLeuThrLysProThrThrThrSerSerGluArgGluArgAspAr 1800	8265 TGGGCCCGTGAACGTCCTGAAAGGGCCTGTG	1800 gAspArgGluArgAspArgAspArgGluArgGluLysSerIleLeuThrSerThrThrTh 1820 ::::::	8296	1820 rvalGluHisAlaProllerrpArgProGlyThrGl 1832	GTGAACGCCACGGTGGGCACAGTGAATGCCGCCCCAGGCACAGT	uGlnSerSerdlySer	AGTGCAGTGACCGTCACAGCGGGTGC	1838SerGlySerSerGlyGlyGlyGlyGlyGlyGlyGlySer	GGTTACTGCTGCATCTGGTGTGTAACGGCCACAAACGGCACGGTGACAATTGGCAGGGGGCACAATTAGGTGACAATTAGGTGTAACGGCCACAAACGGCACAAAAAAAA	1848SerSerArgProAlaSerHisSerHisAlaHisSrFroil 1863
á	3	ò	а	ò	q	ò	qa	ζö	qq	ò	q _Q	ò	đ	λō	đ	ò	đ	ò	q	ò	qa	ζ	qa	ò	qu	ò	QO	λ	ΩD	ò	đ	ò	đ	δ	q	ò

roLysGlnProGlyProValLysLeuGlyGlyGluAlaAlaHisLeuProH 2105 rlleglnGluLeuGluLeuArgSerLeuGlyTyrHisGlySerSerTyrSe, 2049 1yValGluProValSerProValSer-----SerProSerLeuT 2065 ysGlyLeuProLysHisLeuGluGluLeuAspLysSerHisLeuGluGlyG 2085 : CAAACTGCCTACAGAAGTCAACATGTCCCCTCGGGGCCCAGCATCCCAGC 9230 :::: TGTCTCCCCATTTGGCAGCTGCAAAGCTAGATGCTCAT------TC 9278 ----- 9343 ||| |cagcatccaccagagcagtctgtcatcatgccaccccacagcatcaccca 9461 1212 ----ValProProValSerGlyHisAlaThrIleAlaArgThrProAla-- 1999 1999 2013 caaccraagarccacacricacacccaccacac-----ccrccrac 9170 aSerAlaSerAspPro-----HisArgGluLysThrGlnSerLysProPh 2031 CCTGAGCCACCTCTCCCAGGGGGAGGTGAGAATGAACACTCCCACGCTGCC 9521 CCGTCAACAAAGTGCAAACAGAGAGCGAGTGCTAATGAAAACAGTCGGTT 8558 ThrGlnAspAlaLeuGlnGlnArgProSerValLeuHisAsnThrGlyMe 1883
::: :: ||||||||::: ||||
:rccargccrgrGaTcGATCGTCGGCA------GACGCGGC-- 8607 IleThrAlavalGluProSerLysProThrvalLeuArgSerThrSer-- 1902 SerSerprovalArgProAlaAla-----ThrPheproProAlaTh 1917 LeuGlyGlyThrLeuAspGlyValTyrProThrLeuMetGluProValLe 1937 :::||||
------GCAATGGACATTGAGTATCAGCAGTCAAGTCCAAGTCCCA 8768 3GluAlaProArgValAlaArgPro-----GluArgProArgAlaAspTh 1955 :::::: |||:::|||
rdaticigicacagcaicgadgciccarccaaggcccrcaaggrccigc 8828 | :::||| SAACGIGGCCACCCATICCACGITGGIACIGACCGCCCAGACAIAIAAIGC 8888 AhrgserGlyLeuGluProAlaSerSerProSerLysGlySerGluProAr 1982 CTCTCGGTGTCCACGCCTGTCACCCAGGGAGGCACAGTGAAGGTTCTCAC 9002 AAACAAGAAGCTTGCTGACCCCGTCACCCTTAAAATCGAGACCAAGGTCCT 9122 ------LysAsnLeuAlaProHisHisAlaSerProAspProProAl PheLeuAla----

Page 384

919 GilyAapProArghlaanhlaaga-ProChings——ProLeukap— 539 GOTGAG——CATCHAAAACCCCTCTCTCTAACAACCCCTTACCAACACACAC	ThrargvalProSerAspSerAlaileThrTyrArgGlySerIleThr GCAGGTGCACGCGCTGCAGCTTCTGAG	152		1470 LysHisAspValArgSerLeulleGlySerProGlyArgThrPheproProval 1487
949 GIJVABPPEANERAIAARANIASERPROGIDLY9—PPROLEMBP— 959 AGRANGAGUTTGAGGGGAGGCTTCCTTCCLANGAGCGTTCCCTTCCTTCCLANGAGCGTTCCTTCCTTCCTTCCTTCCTTCCTTCCTTCCTTC	8888888	8 & 8 & 8 & 8 & 8	8 6 8 6 8 6 8 6 8	8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8
8 4 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6	949 GlyAspProArgAlaAsnAlaSerProGlnLysProLeuAsp	6169 ACTGCAGCTGGTGGTGGCCCCCAGGGAAAAAGGGAAAAAAGGGAAAAAAGGGAAAAAA	1087 GlyLeuHisAspThralaArgProValLeuProArgProProThrileSerAsnProPro 1087 GlyLeuHisAspThralaArgProValLeuProArgProProThrileSerAsnProPro 1107 ProLeuIleSerAscTCCAGGTTAGCAGTGGCAAATCTGCAAGTCTGAAAAT 1107 ProLeuIleSerSerAlaLySHisProSerValLeuGluArgGlnIleGlyAlaIleSer 1127 GlnGlyMetSerValGlaCrGTCCAGGGGGGGCTGCAGGCAGGCAGGGAAA 1127 GlnGlyMetSerValGlnLeuHisValProTyrSerGluHisAlaLysAlaProValGly 1127 GlnGlyMetSerValGlnLeuHisValProTyrSerGluHisAlaLysAlaProValGly 1127 GlnGlyMetGlyLeuHroLeuHisValProTyrSerGluHisAlaLysAlaProValGly 1147 ProValThrMetGlyLeuProLeuProMetAspPro	1172

CTAGAGAGAAAAGATTCTGACTTCATTTCTGGTAGGATCTATGGGAAGCAGACA LysalabroalaalaalaserGluGluAlaalapheProProValVal	749GluserIleProSer	
4 6 4 6 4 6 4	6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6	8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8
	4000 CATGGTTCCTTCCATGAAGATGAAGATCCCTATAGATGACTAGAGAGAG	623
8 8 8 8 8 8 8 8	8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8	6 H 6 H 6 H 6 H 6 H 6 H 6 H 6 H 6 H 6 H

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cc is afflicted with breast cancer. The method comprises comparing the level of expression of a marker (gene/polypeptide see ACC5076 to ACC50334 and CC ABR47386 to ABR47632) in a patient sample and the normal level of expression of the marker in a control non-breast cancer sample, where a significant increase in the level of expression of the marker in the CC patient sample and the normal level is an indication that the patient is a filitted with breast cancer. The breast cancer associated sequences from the present invention have cytostatic activities and can be used in gene CC therapy. The method is useful for diagnosing and treating breast cancer. N.B. The sequence data for this patent did not form part of the printed specification, but was obtained in electronic format directly from WIPO CC at ftp.wipo.int/pub/published_pct_sequences

XX Sequence 12227 BP; 3448 A; 3248 C; 3069 G; 2462 T; 0 U; 0 Other;
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88888 8 8	therapy. The ment N.B. The sequent specification, at ftp.wipo.int	sthod schod sce da but w '/pub/	34 <u>5</u> 5	did not successions	treating breas orm part of the rmat directly f.	0
S	Sequence	12227 BP; 3	448 A; 3248	C; 3069 G; 2462	T; 0 U; 0 Other;	
Alignmes Pred. No Score: Percent Best Lo Query Ma	gnment Scores: d. No.: re: cent Similarity: t Local Similarity ry Match:		.92e-09 86.00 10.91\$ 9.39\$	Length: Matches: Conservative: Mismatches: Indels:	12227 638 139 1071 1207 150	
0-SD	US-09-522-753-	-5 (1-2517)	x ACC50291 (1	1-12227)	-	
ò	41	GluTyrGlnHi	BHisSerArgAsı	GluTyrGlnHisHisSerArgAspTyrAlaSerHisLeuSerPr	oGlySerllelleG	09
Вр	2077	GACTATAACCA	AGATCGTACATA:	rtatgagagtgttcga	CCAGGCACTTATCCTGA	2136
È	61	ProGlnArgArg-	- 1	rLeuLeuSerGluPhe	-ArgProSerLeuLeuSerGluPheGlnProGlyAsnGluArgSer	79
д	2137	GATTCCAGGCG	GGACTATCCAGC	· TCGAGGGAGAGAGTTT	GATTCCAGGCGGGACTATCCAGCTCGAGGGAGAGAGTTTTATTCAGAATGGGAAACTTAC	2196
È	80	Gln				80
g	2197	CAAGGAGACTA	CTATGAATCACG	ATACTACGATGATCCT	 CAAGGAGACTACTATGAATCACGATACTACGATGATCCTCGGGAATACAGGGATTACAGG	2256
ò	81		-GlubeuHisber	ArgProGluSerHis	GluLeuHisLeuArgProGluSerHisSerTyrLeuProGluLeuGly	96
QQ	2257	AATGATCCTTA	 AATGATCCTTATGAACAAGATATTAGG		::: -GAATATAGTTACAGGCAAAGGGAACGA	2310
à	97	LysSerGluMe	tGluPheIleGlu	LysSerGluMetGluPhelleGluSerLysArgProArg	Leu	110
qq	2311	GAAAGAGAACG	TGAAAGATTTGAC	GTCTGACCGGGACAGA		2370
à	111	GlubeubeuProAspPro-		-LeuLeuArgPro	SerProLeu	123
g	2371	GAACGAAGTCA	аастсстсттса	CTTGCGACGTCCACAG		2430
ò	124	LeuAlaThrGl	LeuAlaThrGlyGlnProAlaGlySerGlu	/SerGlu	AspleuThrLysAspArg	139
В	2431	CAGGCAGAGAG	GTTGCCGAGTGA	ricidadaggaggerr	CAGGCAGAGAGGTTGCCGAGTGATTCTGAGAGGAGGCTTTACAGCCGATCCTCAGACCGG	2490
è	140	SerLeuThrGl	erLeuThrGlyLysLeuGluProValSerProPro	SValSerProPro		151
qq	2491	AGTGG	AAGCTGTAGCTC	ACTCTCCCTCCAAGA	GGAAGCTGTAGCTCTCCCTCCAAGATATGAGAAACTGGACAAGTCT	2544
à	152			oProHisThrAspPro	SerProProHisThrAspProGluLeuGluLeuValProPro	165
g	2545	CGTTTGGAGCG	CTATACAAAAAA	rgaaaagacagataaa		2604
ò	166	ArgleuSerly	BGluGluLeuIle	pAr	ValAspArgGluileThrMet	185
q	2605	AGAGTGGAGAG	AGAGAGACGCTTA	AATACGGAAGGAAAAA		2664
à	186	ValGluGlnGl	ValGluGlnGlnIleSerLysLeu-		ysGlnGlnGlnLeuGluGlu	202
qq	2665	AAGCAGAAACG	CAAAGGAAAGGTT	rcactcccctagttct	AAGCAGAAAGGAAAGGATCACTCCCTAGTTCTCAGTCTTCAGAAACGGACCAA	2724
ò	. 203	GluAlaAlaLy	BProProGluPro	GluhlahlahlysProProGluProGluLysProValSerProProProIleGlu	ProProProlleGlu	220
qq	2725	GAAAATGAGCG	AGAGCAAAGCCCT	rgaaaagcccaggagt		2784

2419 9341 GGACC 2426 rgPrc 2426 rgPrc 2434 isSec 2435 rgPrc 2445 rgPrc 245 rgPrc 2461 rTCC 2462 rGCl 2496 isAll 9701 GTCT ILT 95 ACC50291 stan ACC50291, 12-JUN-2003 Breast cancer Human; breast Human; breast Human; breast Human; breast Human; breast 16-JAN-2003; 21-JUN-2001; 27-JUN-2001;	PR 18-JUL-2001; 2001US-0 PR 25-SEP-2001; 2001US-0 PR 14-MAY-2002; 2002US-0 XX (MILL-) MILLENIUM PHA XX (MILL-) MILLENIUM PHA XX (MILL-) Gannavarap PI Lillie J, Gannavarap PI Bast RC, Hortobagyi XX WPI; 2003-210381/20. DR P-PSDB; ABR47592. XY Breast cancer diagnos PT of a marker in a pati PT cancer sample. XX C The present inventior XX C The present inventior
	2310 lyThrGluIlePheAsnMetProAlaileThrGlyInrGlyEuwerLintlyArtgSect 2330 2310 lyThrGluIlePheAsnMetProAlaileThrGlyInrGlyEuwerLintlyArtgSect 2330 2330 lnalaValGluHis
8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6	8 8 8 8 8 8 8 8 8 8

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|||
| 3ACTICCCAGAGACCCGIGGAIAIGGITCAACTICIGAAGAAGIACCCC 9640
                                                                                                                                                                                                                                                                                                                                                                                                           AlaGlySerGlyPro-----------LeuAlaGlyProHisH 2496
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         osis or treatment by comparing the level of expression tient sample with that in the control non-breast
                                                                     AGGCCACCTGAGGCCTCACACCCAGGTTCAGAGGGGCACAAGGCAGAAAACA 9400
                                                                                                              ------ProSerVal------SerSerValH 2434
                                                                                                                                   spCysAsnArgArgThrProLeu---ThrAsnArgValTrpGluAspA 2453
                                                                                                                                                                                                            |||||||
sacrgecceaaacagecgrrgtrgreecaacaacererggeeceage 9520
                                                                                                                                                                                                                                                          AlaglySerThrPro-----PheproTyrAsnProLeuIleMetArg, 2471
                                                                                                                                                                                                                                                                                    -------GlyValMetAlaSerProProP 2482
                                                                                                                                                                                                                                                                                                                                                                                                                                    ----- AAAGGGGACCCAGACGGGAGTAGAGCAGCCTCGCCTCCCAGCT 9340
                                      ------AlaSerGlyAspA 2426
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 on describes a method for assessing whether a patient
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              pu M, Glatt K, Hoersh S, Kamatkar S;
JE, Myer V, Wang Y, Xu Y, Zhao X, Meyers RE;
GN, Pusztai L, Meric F, Sahin A, Mills GB;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              ; cytostatic; gene therapy; gene; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ated cDNA sequence SEQ ID NO:422.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         3luGluProLysPro 2504
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    SAACGICCIGGCCCA 9726
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    422; 128pp; English.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     DNA; 12227 BP.
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-0301572P.
-0306501P.
-0325002P.
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2119 euLeu	& 	7357	8
	qa	1838SerGlySerSerGlyGlyGlyGlySer	ð :
· •d	8 8	7297 CAATGCCGCTGCGAGTGCAGTGAATGCCACAAGTGCAGTGACCGTCACAGCGGGTGC	q
2085 luLeuArgProLysGlnProGlyProValLysI 	ò £	1832 uGlnSerSerGly8er	ò
 8257 TGCAT	-	1820 rValGluhisAlaProlleTrpArgProGlyThrGl 1832	පි සි
819/ ICCICGACCAAGIGGACCCGGGCCAICCICAFI 2065 hrHisAsplvsGlvLeuProLvsHisLeuGluC	8 8	7214	q
2049 rPro-GluGlyValGluProValSerProValS	ð 1	1800 gAspArgGluArgAspArgAspArgGluArgGluLysSerIleLeuThrSerThrThrTh	ò
AGATC	. 요	1780 yGlyProThrHisLeuThrLysProThrThrSerSerSerGluArgGluArgAspAr 1800	& A
	ð		අ
2013 arioficoAlaseiAlaseiAsprio	∂ 8	1760 rLeuProThrAlaProGlnProPheSerSerArgHisSerSerSerProL	ò
	а 8	7117	đ
	70	1740 uProValLeuValProProThrProGlyThrProAlaThrAlaMetAspArgLeuAlaTy	ò
7981 TGTCACCACAAACAAGAAGCTTGCTGACCCCG	qa	1720 uAlaLeuAsnTyrAlaAlaGlyProArgGlyIleIleAspLeuSerGlnValProHisLe 1740 1073	<u>ک</u> ج
	ò	7034	셤
1985ValProProValSerGlyHi	8	1700 aThrAlaMetAlaGlnArgAlaAspMetLeuArgGlyLeuSerProArgGluSerSerLe	ò
	අධ	1680 uAsnargGlnThrileIleAsnaspTyrileThrSerGlnGlnMetHisHisAsnThrAl 1700 1018 AAATTCAAGAAATA	රි සි
1982 aProleur	8 ò	6369	ପ୍ର
1962 sProProAlaArgSerGlyLeuGluProAlaSe	ð í	1662 OHisLeuTyrProProTyrLeuIleArgGlyTyrProAspThrAlaAlaLeuGl	8
::: AGGCTATGCGAACGTGGC	- qa	1646 aAlaAlaTyrTyrLeuProArgHisLeuAlaProAsnProThrTyrPr 1662	8 8
/68/ GGTCAAACCTGATTCTGTCACAGCATCGCAGCC	8 8 	6839	qq
1937 uLeuProLysGluAlaProArgValAlaArgPr	8	1637.eProArgGly1leProLeuAspAlaAl	ò
	. A	1617 yValSerGlyValAspLeuTyrArgSerHisIleProLeuAlaPheAspProThrSerII 1637	8 8
7585 GCAGAAGACCGAAGGCCCACAGCGGATCAGCGC	ය ද	6764 AGTGTCACTTCCACAAGTGTCACCACAGCCATTGCAGAGCCTG	đ
1903ThrSerSerProValArgProAlaAl	ò	1597 8SerfhrValProGluHisHisProHisProJleSerProTvrGluHisLeuLeuArdGl	8 8
	i 6	1578 LysAlaSerGlnAspArgLysLeuThrSerThrProArgGluileAlaLysSer-ProHi 1597	8 8
7477 CCACCCAGGGTCCATGCCTGTGATCGACGATCC	8 8	::: GCTCAGTCTACTCCATCTC	qa
1863 eSerProArgThrGlnAspAlaLeuGlnGlnA1	8 1	1561 ThrMetArgGluProThrProArgleuGlnGluGlySerLeuSerSerSer	ò
 AGTGATTGCGCCGT	· 8	1541 LeuThrTyrGluAspHisGlyAlaProPheAlaGlyHisLeuProArgGlySerProVal 1560	ò 6
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.	1848SerSerArgProAlaSerHisSerHisAlaHisGlnHisSerProIl
0	7417 AGTGATTGCGCCGTCAACAAAGTGCAAACAGAGGGGAGTGCTAATGAAAACAGTCGGTT 7476
_	uHisAsnThrGlyMe 1
0	7477 CCACCCÁGGGTCCATGCCTGTGATGGATCGTCGGCAGACGCGGGC 7525
`~	1883 tLysGlylleIleThrAlaValGluProSerLysProThrValLeuArgSerThrSer 1902
0	7526 -TCAGGGGCGGGGCTGCGTGAACACTTCTGAAGGGGTTGTGCTCCTGAGTTACTCAGG 7584
`	191
۵	7585 GCAGAAGACCGAAGGCCCACAGCGGATCAGCGCCAAGATCAGCCAGATCCCCCCGGCCAG 7644
_	1917 rHisCysProLeuGlyGlyThrLeuAspGlyValTyrProThrLeuMetGluProValLe 1937
0	7645 TGCAATGACATTGAATTTCAGCAGTCAGTCCAAGTCCCA 7686
5.	937 uLeuProLygGluAlaProArgValAlaArgProGluArgProArgAlaAspTh 195
0	7687 GGTCAAACCTGATTCTGTCACAGCATCGCAGCCTCCAAAGGCCCTCAAGCTCCTGC 7746
~	1955 rGlyHisAlaPheLeuAlaLy 1962
0	7747 AGGCTATGCGAACGTGGCCACCCATTCCACGTTGGTACTGACCGCCCAGACATATAATGC 7806
.	1962 sProProAlaArgSerGlyLeuGluProAlaSerSerProSerLysGlySerGluProAr 1982
0	7807 CTCTCCTGTGATTTCGTCTGTGAAGGCCGATAGGCCATCCTTGGAGAGCCCGA 7860
.	1982 gProLeu 1984
0	7861 GCCCATTCACCTCTCGGTGTCCACGCCTGTCACCCCAGGGAGGCACAGTGAAGGTTCTCAC 7920
.	1985ValProProValSerGlyHisAlaThrIleAlaArgThrProAla 1999
0	PACACACCCCTGTGCTGGTTCACAACCAGCTGGTCCTCACCCCAAG
.	199
0	AACAAGAAGCTTGCTGACCCCGTCACCCTTAAAATCGAGACCAAGGTCCT 804
.	2000LysAsnLeuAlaProHisHisAlaSerProAspProProAl 2013
0	808
<u>~</u>	2013 aProproAlaSerAlaSerAspProHisArgGluLysThrGlnSerLysProPh 2031
0	CAGC 814
.	2031 eSerlleGlnGluLeuGluLeuArgSerLeuGlyTyrHisGlySerSerTyrSe 2049
0	TC 819
<u> </u>	rPro-GluGlyValGluProValSerProValSerSerProSerLeuT 206
0	arccicariccaagggaagccacccca
5.	yLeuProLysHisLeuGluGluLeuAspLysSerHisLeuGluGl
0	
.	2085 luLeuArgProLysGlnProGlyProValLysLeuGlyGlyGluAlaAlaHisLeuProH 2105
o [*]	caccaaceccacaccarecreecrecaeécareccaerecéca 831
`	211
0	CATCCACCCAGAGCAGTCTGTCATGCCACCCCCACAGCATCACCCA 837
`~	2119 euLeuG 2121

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		962	GGTGAC	963EquiyaGinLequiyaGinArgalaAlaAlatieftoFolicatiosolisation 773		# # # # # # # # # # # # # # # # # # #		AAGGAACCTGCAGAAACACTCAAGCCACCTGAGGGATGGCGGTCGCCAAGGTCCCAAGAAAA				1027 AlaProProAlaAspLysGluAlaPheAlaAlaGluAlaGluLysLeuProGlyAspPro 1046		ProCysTrpThrSerGlyLeuEroPheFroValProFroArgGiluvallteLySAlassel	GCTGGCACAGACAAAAACCCCCCCCGGGGGGGGCCCCCGGGGGG	106 Frontabalertomepriocentarization		5354 GGAAGATCTCGAAACTCCCAGGTTAGCAGTGGACAATCTGCAAGTCTGAAAAT 5407		5408GIGGAIGCIGCIGICAGICCCAGGGGGGCIGCAGGGCAGGGGAGGGGAA 5461		SAGAGTCCCCAAAAGGAGGATGGT	1147 ProvalThimetGlyLeuProLeuProMetAspPro		 	1172	5639 CAGGCCGTGGAACACATCGCAAAGCTCGCTGAGGCCTCTGCCTCTGCTGCTATAAGGCA 5698		5699 GATGCACCAGAGGGCTTGCCCCAGAGGACAGGGACAAGCCTGCACACCAAGCA 5752	1192 AlaSerValLeuArgGlyThrAlaLeuGlySerVal		1204
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1528 ValileValProGluLeuGlyLysProArgGlnSerPro----------------- 1540 1450 SerlleThrGlnGlyThrProLeuLysTyrAspThrGlyAlaSerThrThrGlySerLys 1469 1470 LysHisAspVajArgSerLeujjeGlySerProGlyArgThrPhePro-----ProVal 1487 1488 HisproleuAspValMetAlaAspAlaArgAlaLeuGluArgAlaCysTyrGluGluSer 1507 1508 LeulysSerArgProGlyThrAlaSerSerGlyGlySerIleAlaArgGlyAlaPro 1527 ------Achdeaaageneneacereteseerendige----- 6583 6272 GAAAAGCAGAGTGAGAAACCCCATTCCACTCCTCCTCAGTCATGTACTTCTGACCTAAGC 6331 1395 GlualaTyrLysThrGlnAlaLeuGlyProLeuLysLeuLysProAlaHisGluGlyLeu 1414 1415 ValAlaThrValLysGluAlaGlyArgSerIleHisGluIleProArgGluGluLeuArg 1434 1298 SerGlyProProHisGluThrAlaAlaProLysArgThrTyrAspMetMetGluGlyArg 1317 1318 ValGlyArgAlaIleSerSerAlaSerIleGluGlyLeuMetGlyArgAlaIleProPro 1337 1338 GluarghisserpronishisLeuLysGluGlnhisHisIleArgGly-----Ser 1354 1355 IleThrGlnGlylleProArgSerTyrValGluAlgluAlgGluAspTyrLeuArgArgGlu 1374 6227 GCAAATGAGGGGACAACAGTACAGCACCCCGAAAGCCCCACAG-------GAA 6271 1375 AlalysleuLeuLysArgGluGlyThrProProProProProProSerArgAspLeuThr 1394 5963 TCTTCTAGGCCTCCAGTCAATGCTCCTGACCCCTCAGCCGGCCCAACAGATACCAAGGAA 6022 1258 GlyargGluAspSerLeuProLysGlyHisVallleTyrGluGlyLysLysGlyHisVal 1277 6023 GCCAGAGGAAATAGCAGTGAAACCTCACACTCAGTGCCAGAAGCCAAAGGGTCTAAA--- 6079 1278 LeuSerTyrGluGlyGlyMetSerValThrGlnCysSerLyygGluAspGlyArgSerSer 1297 6080 ------GAAGTGGAAGTCACTCTTGTTCGGAAAGACAAAGGGCGCCAG--- 6121 1215 ThrargVal-------ProSerAspSerAlalleThrTyrArgGlySerlleThr 1230 1231 HisGlyThrProAlaAspValLeuTyrLysGlyThrIle---ThrArglleIleGlyGlu 1249-ProSerArgLeuAspArg 1257 5813 GAAAACTICCCAGCACCICCACCITAICCIGGAGAAICCCAGACAGAICTGCAACCCCCC 5872 6332 AAGATTCCCTCCACAGAG--------...----GTGGCTCCTGTA 5873 GCAGGTGCACAGGCGCTGCAGCCTTCTGAG----1250 AspSer----6584 6122

whether the comparison indicates a likelihood disease). (M1) is useful for identifying modulators of the p53 pathway. A probe for HM expression is useful for diagnosing breast, colon, kidney, lung and ovarian cancer, in a patient, where the cancer has greater than 25 % expression level. Color and the patient of the patient of the cancer has greater than 25 % expression level. Color and cancer, and the p53 pathway, such as, angiogenesis is related the cancer in the p53 pathway, such as, angiogenesis, apoptotic or cell proliferation disorders (e.g. cancer). Another two new methods (M2 and M3) are useful for modulating the p53 pathway of a cell, thus restoring the p53 function of the cell, so that the cell undergoes normal to proliferation or progression through the cell cycle. (M2) and (M3) are useful for treating defects in the p53 pathway such as angiogenic, also useful for retaing defects in the p53 pathway such as angiogenic, apoptotic or cell proliferation disorders. The present sequence is an HM uncleic acid encoding a p53 pathway modifying protein

Sequence 11145 BP; 3179 A; 2966 C; 2736 G; 2264 T; 0 U; 0 Other;

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						-				Leu	CGATT	-SerProLeu	CCTCT	spArg	Acced	:	AGTCT	roPro	 CGTTTGGAGCGCTATACAAAAATGAAAAGACAGATAAAGAACGAAC	
		GlutyrGlnHisHisSerArgAspTyrAlaSerHisLeuSerProGlySerlleIleGln	GACTATAACCAAGATCGTACATATTATGAGAGTGTTCGAACTCCAGGCACTTATCCTGAG	ArgProSerLeuLeuSerGluPheGlnProGlyAsnGluArgSer	GATTCCAGGCGGGACTATCCAGCTCGAGGGAGAGATTTTATTCAGAATGGGAAACTTAC		 Caaggagactactatgaatcacgatactacgatgatcctcgggaatacagggttacgg	GluLeuHisLeuArgProGluSerHisSerTyrLeuProGluLeuGly	GAATATAGTTACAGGCAAAGGGAACGA	!	:::	SerP	 gaacgaagtcaaagtcctgttcacttgcgacgtccacagagtcctggaggcgtctct	-AspLeuThrLysAspArg	 CAGGCAGAGGTTGCCGAGTGATTCTGAGAGGAGGCTTTACAGCCGATCCTCAGACCGG		-dgaagctgtagctcactcccctccaagatatgagaaactggacaagtct	SerProProHisThrAspProGluLeuGluLeuValProPro	TTGAT	
		3G1ySe	AGGCAC	эд1удв	AGAATG		ATACAG	rLeuPr	CAGGC		TGAGAC		TGGAG	pLeuTl	CCGAT	1	GAAAC	uGluL	BAACTT	
11145 638 379 1071 1207		SerPr	ACTCC	GlnPr	TATTC		CGGGA	SerTy	ragita		AGACCA		BAGTCC	A8	TTACAG		ATATGA	ogluLe	AGAACO	
ive:		HisLeu	STTCGA	31uPhe	GAGTTT		GATCCI	SerHie	GAATA	ProArg	GACAG	Pro	CCACA	1 1 1	AGGCT	Pro	ccaag	Asper	GATAA	
Length: Matches: Conservative Mismatches: Indels: Gaps:	45)	laser	AGAĞT	euSer	GGAGA	:	PACGAT	roglu	- 1	ysArg	3ACCGG	LeuLeuArgPro	cgacgi	31u	GAGAGG	SerPro	1000 1000 1000 1000	HisTh	AAGACZ	
Lengt Match Conse Misma Indel Gaps:	(1-11145)	SPTyrA	ATTATG	erLeuI	CTCGAC		GATAC	euArgi	TTAGG	luseri	AGTCT	Leu	ACTTG	lyser'	ATTCT	roval	CACTC	ProPro	AATGAA	
o		rArgA	TACAT	gPros	TCCAG		ATCAC	uHi BL	: VAGATA	neIleG	ATTTE	6-	CTGTTC	roAlaG	CGAGTC	euGluE	GTAGCT	Ser	CAAAA	
1.79e-09 586.00 30.91% 19.39% 7.43%	x ACD13385	внівЅе	AGATCO	gA1	GGACTA		CTATG	-Glufe	 - AATGATCCTTATGAACAAGATATTAGG	LysSerGluMetGluPheIleGluSerLysArgProArg-	TGAAA	GluLeuLeuProAspPro	AAGTC	LeuAlaThrGlyGlnProAlaGlySerGlu	GTTGC	SerLeuThrGlyLysLeuGluProValSerProPro-	SAAGCT	1	SCTATA	
 456447	(1-2517)	GlnHi	PACCA	ArgAr	AGGCG		GACTA		rcctta	rG1uMe	AGAACG	uLeuPr	AAGTC?	aThrG]	- AGAGAC	uThrG]			GGAGC	
cores: iilarity: Similarity ::	5 (1-2	31uTyr	SACTAT	ProGlnArgArg	GATTCC	Gln	CAAGG		AATGAT	LysSeı	GAAAG	gluLe	GAACG	LeuAl	CAGGC	Serle	AGT		CGTTT	,
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|||::: :::||| ::: ||| 1523 AGAGTGGAGAGAGAGAGGCTTAATACGGAAGGAAAAGTGGAAAAGGACAAAACTGAC 1582

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166 ArgleuSerLysGluGluLeulleGlnAsnMetAspArgValAspArgGluIleThrMet 185

1583 AAGCAGAAACGGCAAAGGTTCACTCCCCTAGTTCTCAGTCTTCAGAAACGGACCAA 1642

GlualaalaiysProProGluProGluLysProValSerProProProIleGlu-----

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186 ValGluGlnGlnIleSerLysLeu-----LysLysLysGlnGlnGlnLeuGluGlu 202

1643 GAAAATGAGCGAGAGCAAAGCCCTGAAAAGCCCAGGAGTTGTAATAAACTGAG		-
221		232
233		249
1763 ACTCGAGTGAAAAAAAAAAAAAAAAAAAAAAAAAAAAAA		9 8
262		280
		290
291		306 2056
307 GlualaLeuGluLysLysValGluArgIleGluAsnAsnProArgArgArgAlaLysGlu 		326 2092
327 SerLysValArgGluTyrTyrGluLysGlnPheProGluIleArgLysGlnArgGluLeu		346 2134
. G	SerMetSerAlaAla GAGATGGAGGATGCT	366 2158
ArgSerglu 	SerGluIleIleAsp ::: ATTCTGCTGAGGGAA	377 2218
ω σ	GAATCTAAAAAATC	389
	AlaAspGlnGlnArg AAAGACTGTCAGGAG	406 2326
407	ValtyrLys acctacaagcaagac	424
425AspArgGlnValMetA ::: 385 TGGGAGAACTAGCAGGTGA	GluLysGluThrPhe ::: :caaaaaaGCCCATTC	440
- 5	SerPheLeuGluArg ::: GACCAGAGAGAGAAG	460 2504
461LysThrValAlaGluCysValLeuTyrTyrTyr	CGTAAATCAGGCCAAG	471 2564
472	sSerLeuValArgArg ::: \TCGATCACGCAGGA	485
486	nGlnGlnGlnGlnGln ::: 3GAAATGGAAATAGCC	505

Qy 2191GlyAlaProAlaArgGlySerProHisSerGluGlyGlyLy8ArgSerProGlubroA 2210 Db 8889 TTTCCCAGCAGGACATT-CCTCATGGGGGAGGCAGACCCCCCG 8937 Qy 2210 snLysThrSerValLeuGlyGlyGlyGluAspGlyIleGluProValSerPro 2227 Db 8938TCTGTCCTCGGTGACGGCTGAGGCTGCCCCCCCCCC	RESULT 94 ACD13385 standard; CDNA; 11145 BP. X ACD13385; ACD13385; XX ACD13385; XX ACD13385; XX ACD13385; XX ACD13385; XX Human; SS; Gene; p53 modifier; SEQ ID 56. XX Human; SS; Gene; p53 modifier; cytostatic; cancer; cytostatic; XX Human; SS; Gene; p53 modifier; cytostatic; cancer; cytostatic; XX Human; SS; Gene; p53 modifier; cytostatic; cancer; cytostatic; XX Human; SS; Gene; p53 modifier; cytostatic; cancer; cytostatic; XX Human; SS; Gene; p53 modifier; XX Homo sapiens. XX MC200299122-A1. XX CCCC2001; 2001US-023605EP. XX CX EXEL-) EXELIXIS INC. XX EXEL-) EXELIXIS INC. XX EXEL-) EXELIXIS INC. XX MPI; 2003-156659/15. XX MPI; 2003-156659/15. XX T dentifying modulators of the p53 pathway for use in treating apoptotic YY T dentifying modulators of the p53 pathway for use in treating apoptotic YY T dentifying modulators of the p53 pathway for use in treating apoptotic YY T modulate activity of a human ortholog of genes that modify the p53 YY Pathway in Drosophila. XX The invention relates to identifying (MJ) a candidate p53 pathway in pathway in Drosophila. XX The invention relates to identifying (MJ) a candidate p53 pathway of modulating agent. by contacting an assay system comprising a teference CC for the presence of the test agent, the system provides a reference CC for the presence of the test agent, the system provides a reference CC for the presence of the test agent, the system provides a reference CC activity and ace exide with a test agent under conditions, where but CC activity are activity of the test agent, where provides a reference CC activity are activity of the test agent, where provides a reference	Contacting a cell defective in 53 function with a candidate modulator that specifically binds to a HM polypeptide comprising an HM amino acid sequence, where p53 function is restored), modulating (M3) a p53 pathway CC in a mammalian cell (comprising contacting the cell with an agent that CC specifically binds an HM polypeptide or nucleic acid) and diagnosing (M4) CC a disease in a patient (comprising: (a) obtaining a biological sample CC from the patient; (b) contacting the sample with a probe for HM CC expression; (c) comparing the results with a control; and (d) determining
laLeuGlnGlnArgProSerValLeuHisAsnThrGlyMetLysGlyIleIleThralaV ::: ::: ::: :::	1947	GCACATTCCCATGGGGCCCTGTTTTTTCCCCCATGGGGCCCTGGTGGGGGCCTGGTGGGGGGGG

è	1188 rAlaGlnGluAlaSerValLeuArgGlyThrAla	ź	
; <u>d</u>		ìò	1528 lileValProGluLeuGlyL
ò	SerAspSe	섬	7359 CGTGGAGCCCGCGCCGTTC
Ор		ζO	1548 aProPheAlaGlyHisLeuP
ò	SerIleThrHisGlyThrProAlaAspValLeuTyrLysGl	qa	7419 TGAGGCTGAAGGCCCCC
ΟD		λō	1568 gLeuGlnGluGlySerLeuS
λ	eThrArg11e11eGlyGluAspSerProSerArgLeuAspArgGlyArgGl :	qq	7474AACACTGTGG
qq		ò	1588 rProArgGluIleAlaLysS
λõ	1260 uAspSerLeuProLysGlyHisValIleTyrGluGlyLysLysGlyHisValLeuSerTy 1280	ପ୍ର	7512 CCCCGAGGACGACACTGAGG
qq		ò	1608 eSerProTyrGluHisLeuI
ò	P.	đ	7563
qq		ò	1628 eProLeuAlaPheAspPro7
λō		셤	7564
q		ò	1648 aTyrTyrLeuProArgHisI
δλ	1320 gAlaIleSerSerAlaSerIleGluGlyLeuMetGlyArgAlaIleProProGluArgH1 1340	q	7594CCAAAACCC
Ωp		ò	1668 rLeulleArgGlyTyrPro
λö	1340 sSerProHisHisLeuLysGluGlnHisHisIleArgGlySerII 1355	qa	7630GTGGAGGAGATCCCT
QQ		ò	1688 pTyrlleThrSerGlnGln
δ	1355 eThrGlnGlyIleProArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGluAl 1375	q	7645CAGCGC
QC		ò	1708 spMetLeuArgGlyLeuSe
ò		ପ୍ପ	7692 GCAGGGCCCGCCCCTCO
ф		ò	1728 roArgGlyIleIleAspLe
λō	1395 uAlaTyrLysThrClnAlaLeuGlyProLeuLysLeuLys	셤	7752 CAAGG
g G		λō	1748 roGlyThrProAlaThrAl
λ	1409 -ProAlaHisGluGlyLeuValAlaThrValLysGluAlaGlyArgSerIleHisGluIl 1428	අු	7788 CCAGCATCCGCGCAAACGC
qq		۸۵	1763 hrAlaProGlnProPheSe
δ	1428 eProArgGluGluLeuArgHisThrProGluLeuProLeuAlaProAr 1444	qa	7848 CACGTCCACGCAGCAGACG
qq	7023 CCCCGAGGACGACCTGGACCTGGGGCCCTTCTCCCTGCCGGAGCTTCCCCTGCAGACTAA 7082	ò	1775SerProLeuSerProGl
ò	1444 gProLeuLysGluGlySerIleThrGlnGlyThrProLeuLysTyrAsp 1460	අධ	7908 CATCAAGCTGGATGCCATC
qq	7083 AGATGCCGCAGATGGTGAAGCGGAACCCGTGGAAGAAAGTCTTGCTCCTCCAGAAGAGAT 7142	ò	1794 erGluArgGluArgAspAr
ò	1461 ThrGlyAlaSerThrThrGlySerLyBLyBHisAspValArgSerLeu 1476	qa	:: 1968 CCTGCAGATCAGGAAGAAG
କୁ	7143 GCCTCCAGGGCCCCCCGGGGTCATAAACGGTGGGGATGTTTCCACCGTAGTGGC 7196	ò	1810 rgGluLysSerIleLeuTh
δλ	1477 IleGlySerProGlyArgThrPheProProValHisProLeuAspValMetAlaAsp 1495	ୁ ଶ୍ର	8028 GGCGCCCAGTGCTACGC
අු	7197 TGAGGAGCCGCCGGCACTGCCTCCTGACCAGGCCTCCACCCGGCTCCTGCAGAGCTCGA 7230	δ	1830 lyThrGluGlnSerSerG
ò	1496 AlaArgAlaLeuGluArgAlaCysTyrGluGluSerLeuLysSerArgFroGlyThrAta 1515	ପ୍ର	8088 GCCGCTCAGCAAGCTCCAC
qa	GCCTGAGCCCTCAGGGGAGCCAAAGCTGGACGTGGCTCTAGA	à	1850 rgProAlaSerHisSerHi
λō	1516 SerSerSerGlyGly-SerIleAlaArgGlyAlaProva 1528	_	

31yserserGlyserserGlyGlyGlyGlyGlySerserserA 1850 HisAlaHisGlnHisSerProlleSerProArgThrGlnAspA 1870 rcaagccciaccacagcgaccaaccccracrrcgaara 7967 ArgAspArgGluArgAspArg-----AspArgGluA 1810 ThrserThrThrValGluHisAlaProIleTrpArgProG 1830 ||||||||::: acarccccgrcgrcgccccccccccccccc 8134 gecegaladraticaldadacecrececearceresec 7907 31yGlyProThrHisLeuThrLysProThrThrThrSerSerS 1794 kancakaaaaacacakanccreracreratakakaka 8027 erProArgGluSerSerLeuAlaLeuAsnTyrAlaAlaGlyP 1728 euSerGlnValproHisLeuproValLeuValProproThrp 1748 .jaMet-----AspArgLeuAlaTyrLeuProT 1763 ccctttcaccctccacccagcacctccaccaccaccaccacraa 7847 oThrSerlleproArgGlyrlleProLeuAspAlaAlaAl 1648 sLeualaProAsnProThrTyrProHisLeuTyrProProTy 1668 CACGGCCGAAGCCCCCCCGA----- 7629 oAspThrAlaAlaLeuGluAsnArgGlnThrIleIleAsnAs 1688 nmetHisHisAsnThrAlaThrAlaMetAlaGln-ArgAlaA 1708 ::: |||| | CGAGAAGAGTGCGCCCCACCCCTGCCCCGGTCACCAGGGC 7751 LeuArgGlyValSerGlyValAspLeuTyrArgSerHisIl 1628 sserProHisSerThrValProGluHisHisProHisProIl 1608 39CCTCCCGTGCCGCCCCAGCCGAAGGCCCT----- 7563 ccccchgaacadccccchtagggagcggagacgaadacaggggg 7418 ProArgGlySerProValThrMetArgGluProThrProAr 1568 SerSerSerLysAlaSerGlnAspArgLysLeuThrSerTh 1588 :::: :::::|||::: 3GCACAAGCTCAGGCCGCAGACGGT------GCCGG 7511 ACGIGCCGGAAGAGAGGCCCGIGGGGATCCGGACTCCAG 7358 LysproArgGlnSerproLeuThrTyrGluAspHisGlXAl serSerArgHisSerSer-----

qq	4486 CITGAAAAGTTGGAAGAAGAGGCTCTCCATGAGTACAGAAGAAGACTCCAACGATAGAATC 4545		רמיי סממ
ò	565 AlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgArgLysGly 580	; E	
qq	4546 AGCGAGGICTCCTCTGACAGCTTCACGGACCGAGGAGCCGGGGCTGACTGCCTTC 4605	G (WEST OFFICE
ò	581ArglleThrArgSerMetAlaAsnGluAlaAsnSerGluGluAla1leThr 597	Š †	
Q	4606 CTGGAGGTCTCTTTCACGGAGCCACCTGGAGGACGAAGCCGAGGAGGAGGGCCTGCCT	on d	
ò	598 ProGlnGlnSerhlaGluLeuAlaSerMetGluLeuAsnGluSerSerArgTrpThrGlu 617	Š †	922 erA
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È	618 GluGluMetGluThrAlaLysLysGlyLeuLeuGlu-HisGlyArgAsnTrpSerAlaIl 637	Š 1	939
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ò	637 eAlaArgMetValGlySerLysThrValSerGlnCysLys	Š í	
οp	4786 AAGGAGGGGGGAAGGAACTCCGGCCAGTACGAAAAGGACTTCCTGGAGGGT 4845	gg (-32 9976
δλ	651AsnPheTyrPheAsnTyrLysLysArgGlnAsnLeuAspGlulleLe 666	Ši i	970 LaA
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ò	666 uGlnGlnHisLysLeuLysMetGluLysGluArgAsnAlaArgArgLy 682	ð 1	
qq	4905 TGAATTGTTTCTACCGAAAAGAAAGATAAAAATGATTCCGAGAGAGA	q	5862 CGC
ò	682 gLysLysLyslygAlaProAlaAlaSerGluGluAlaAlaPheProProValValGluAs 702	ờ	
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ò	SysSerGlvProAlaThrVa	à	1019 roA
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łè	AsnasnserSerasnThrG] SerT eProSerProHisThrG alasavathrg	ò	1038 luA
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g	CTCGGCGATGCCAAACTGAAGGAGAAATTCAAGGACGGT-GCAGAGAAAGAAAAAAGGGCGA	ò	1092 Ala
රි :	rProAlaSerGluAlaThrGlyAlaProThrProProPro	අු	::: 6279 TCC
Q C	CCCAGTGAAGATGAGCAACGGGAATGATAAGGTAGCGCCATCCAAAGACCCCAGGCAAGAA	ò	1112 Ala
Ġ	811AlaProProSerProSerAlaProProProValValProLysGluGluLysGluGl 829	qq	6309 AAA
d d	5337 AGACGCCAGGCCCAGGGA-GAAGCTCCTGGGGGACGGCGACCTGATGATGACCAGCTTCG 5395	<u></u>	1131 Val
ò	829 uGluThrAlaAlaAlaProProValGluGluGluGluGluGlnLysProProAlaAlaGl 849	÷ €	
đ	5396 AGAGGATGCTGTCCCAGAAGGACCTGGAGATCGAGGAGCGCCACAAGCGGCACA 5449	3 8	
ò	849 uGluLeuAlaValAspThrGlyLysAlaGluGluProValLysSerGluCysThrGluGl 869	Š €	150 0011
Q	5450 AGGAGAGGATGAAGCAAATGGAGAAAGCTGAGGCACCGGTCCGG 5492	3 8	
à	869 uAlaGluGluGlyProAlaLysGlyLysAspAlaGluAlaAlaGluAlaThrAlaGluGl	Š :	
q	5493	Q	6474 CAC

ProThrLys------ ProAlaProProAlaProProProProProPro- 1002 GlnAsnLeuGlnProGluSerAspAlaProGlnGln-----ProGlySerSerP 1019 ArgGlyLysSerArgSerProAlaProProAlaAspLysGluAlaPhe---AlaAlaG 1038 AlaGlnLysLeuProGlyAspProProCysTrpThrSerGlyLeuProPheProValP 1058 GCTACGACTCTCCCATGCCACCCTCGA---TGGAAGACAGGGGCGCCCTGCCCCGGGT 6158 ProArgGluValIleLysAlaSerProHisAlaProAspProSerAlaPheSerTyrA 1078 IGInLeuHisValProTyrSerGluHisAlaLysAlaProValGly-ProValThrMe 1150 ------cccegaccregacaccrccaagaaccaggacaggacgccar 6413 .lyLeuProLeuPro-----MetAapProLysLysLeuAlaProPheSerGlyValLy 1168 TCCCCCCGGAGCCCAGCTACCTGGAGCGCTGGAGGGTCCCTTCAGCGCCGTCAT 6473 -------ACCCCTACGTCGTGTATC-CTGCCCCAGCTACGAGGAGGTGATG 5802 CACGCCCAGGACCCCGTCCTGCAGCGCCCGATCACTACGCGGACCTCGTGT-TCGACTG 5861 |||||||::
ccaactcgcagcactccacgccgtgcccacgctcccaccagcgcctgctcccctc 5921 1002 |||||||| |CCGCGGAGAAGTTTGCCTGCTTGTCGCCAGGGTACTACTCCCCCAGACTATGGCCTCCC 6218 ProPro-----HisAspThr 1091 aArgProValLeuProArgProProThrIleSerAsnProProProLeuIleSerSer 1111 aLysHisProSerValLeuGluArgGlnIleGlyAlaIle----SerGlnGlyMetSer 1130 IngluGinLeuSerProArgGlyGlnAlaGlyProProGluSerLeuGlyValProTh 1188 aagccggcagacgacgggaggaagggricrgartccrgcraagaaacggcgg 5575 AlaAspGluValAspGluAlaGluGlyGlyAspLysAsnArgLeuLeu----- 938 ------SerProArgProSerLeuLeuThrProThrGlyAspP 951 ArgalaasnalaserproglnLysProLeuaspLeuLysGlnLeuLys---GlnArga 970 AlaalalleProprolleGlnValThrLysValHisGluProProArgGluAspAlaA 990 laLeuLysAlaGluLysLysGluGlyGlySerGlyArgAlaThrThrAlaLysSerSe 909 lyalapro------Ginaspseraspseralainroys-S :||| |CAGAGGGCGTCTTCTCAAG------

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                                                                                                                                                                                                                                                                                                                                                                                                                                                    AlaThrGlyGlnProAlaGlySerGluAspLeuThrLysAspArgSerLeuThrGlyLys 144
                                                                                                                   The present invention relates to human tumour suppressor gene 16 (TSG16; see AAH23688). The present sequence is a partial TSG16 genomic sequence. TSG16 was isolated from chromosome 16q24.3. TSG16 suppresses cellular proliferation. TSG16 is useful for treating disorders associated with decreased expression or activity of TSG16, e.g. cancers, (auto)immune disorders, inflammation, complications of wound healing and infections (by viruses, bacteria, fungi, parasites, protozoa or helminths)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    MetValGluGlnGlnIleSerLysLeuLysLysLysGlnGlnGlnLeuGluGluGluAla
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               LeuGluGlyLeuGlyProGlnValGluLeuProLeuTyrAsnGlnProSerAspThrArg
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----TTTATAGGAAGATTCTGCTTTTG
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3149 ACGATCAGCGAGATCGGTGTTTTCTGACTTGTCCGATT-----CATCCTTTGATT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         -----SerGluMetGluPheIleGluSerLysArgProArgLeuGlu-----
                 ۵,
                  Crawford
                                                         nucleic acid representing the human tumor suppressor gene TS
ful e.g. for diagnosis and treatment of tumors, inflammatory
                                                                                                                                                                                                    Sequence 9115 BP; 2368 A; 2674 C; 2571 G; 1502 T; 0 U; 0 Other;
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                  Kochetkova M,
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WOMEN'S & CHILDREN'S HOSPITAL
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4485 4425 4365 546 4035 489 547 LysGluLysThrAspAspThrSerGlyGluAsp-----AsnAspGluLysGluAlaVal 564 3870 395 435 LysargargasnHisalaargLysGlnTrpLysGlnLysPheCysGlnArgTyrAspGln 304 265 GlnTyxHisGluAsnIleLysIleAsnGlnAlaMetArgLysLysLeuIleLeuTyrPhe 284 GlnProMetProArgSerSerGlnGluGluLygAspGlu------LysGluLygGlu ::: ::: |||||||| 1366 AAGGCTGAAAAGAAGCTGCCGAAGAGCTAAAAGCTAAAGTGAAGTCG LysGluhlaGluLysGluGluLysProGluValGluAsnAspLysGluAspLeuLeu GACAAAGAACATTCAAAGGAGAGGAGTCCTCGAGAAGTGCCGACGCGGAAAAAGCCCTG 454 IlealaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyrLeuThr ---AspGlyLeuSerGluGluGluAsnLeuGluLysGlnMetArgGlnLeuAlaValIle ProProMetLeuTyrAspAlaAspGlnGlnArgIleLysPheIleAsnMetAsnGlyLeu GlutysgluThrPheArgGluLysPhe------Met ---GluAsnTyrLysSerLeuValArgArgSerTyrArgArg GCAGACATCTTCACAGATGAGAGTGAGGAC-------GACAGAGACAGCTGC MetAlaAspProMetLysValTyrLysAspArgGlnValMetAsnMetTrpSerGluGln -----LysAsnPheGlyLeu ---AAGGAGTGTGGCTGCGAGAGTGGCTTCAAGGACAAGTCCGACGGCGACTTTGGGAAG LeuMetGluAlaLeuGlu------LysLysValGluArgIleGluAsnAsn ProArgArgAlaLysGluSerLysValArgGluTyrTyrGluLysGlnPheProGlu 340 IleArgiysGlnArgGluLeuGlnGluArgMetGln--------SerArgValGlyGln 1811 TTTAAAGAGAAAAAGATACCAAGGAAAAACATAAAGACACACATGGCAAAGACAAAGAA ArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluHisGluValSerGluIleIle ::: :::::: ||||::: |||3751 GACAAAGACAAAGATTTGATAAATGT |||| :::::::||| |---GCCAAGCCCGAGGAGGAGCTG 446 GlnHisPro------ГувгувАвп------

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q	5159 CAGCAGGGGAAGCAAACGTGTCCTTTCACCAGCTGGCAGCCTTGAACTTACCATGGAAA 5218	ò	1586
λō	MetG	qq	6116 ATGTATA
q		ò	1594Lys
6		අු	::: 6176 GCAAGAA
qq		λo	1612 uHisLeu
λo	ysGluGlnHisHisIleArgGlySerIleThrGlnGlyIleProA	qq	6216 -CACTG-
qq	•	λō	1632 eAspPro
δ		Ω	6241 AAGCACT
q		δ	1649 rTyrLeu
δ		ηqα	6299
qq	ACCCTGCATTGTCCC	λo	1669 uIleArg
δ		q	6349 GCCGAAG
g		λõ	1684
ò	/alLysGluAlaGlyArgSerIleHisGluIleProA	qa	6409 GCTGTGG
đ	AGCGTGAAGA	ζ	1699 rAlaThr
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ф		λΌ	1736 nValPro
ò		qa	6565 GACTCC
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à		qa	 6625 ACACGA(
qq		δ.	1769 rSerArg
ò		qu	6685 AGCCAG
q	TGTACAACC	λŏ	1789 hrThrT
ò		qq	6738
ΩĐ	aaagctgctttgtccctcctgaggtctaagcagaag	λ	1809 luArgG
à		qa	6742 AAAGAC
d d		Võ	1829 ProGly
ò	nrTyrGluAspHisGlyAlaProP	q ₀	6799 CCAAGC
qq	5918 CATCCAGCTCCCGCAAGCCCCGCATGACAGAGGTT	ò	1849 SerArg
'n		q	6820 CGTCCA
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ò	/alThrMetArgGluProThrProArgLeuG	qa	6880 TCCCCG
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λ	1570 ln-GluGlySerLeuSerSerLysAlaSerGlnAspArg-LysLeu 1585	ନ୍ଧ	6931
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1586ThrSerThrProArgGlulleAla	1593
	GCT 6175
94LysSerProHisSerThrValProGluHisHisProHisProIleSerProTyrGl	rGl 1612 6215
SerGlyValAspLeuTyrArgSerHisIleProLeuAlaPh	163
6216 -CACTG-ACGTCCGGC	624
1632 eAspProThrSerIleProArgGlyIleProLeuAspAlaAlaAlaTy ::::::	aTy 1649
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6299AGGCCCACAGCAAAAAGTGCCAAGAGACAGGGGTGCTGGAGGAGCTTGGAA	45.4
1669 ulleargGlyTyrProAspThrAlaAlaLeuGluAsnArgGln	AGAG 6408
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6409 GCTGTGGAGGAGCACCAGTTTTCGGACCTGGAGGÁCTCGGACTCAGACTCAGACCTGGAC	3GAC 6468
1699 ralaThrAlaMetalaGlnArgAlaAspMetLeuArgGlyLeuSerProAr 	Ar 1716 AGAG 6528
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1736 nValProHis	hrPr 1751 GGCC 6624
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685 AGCCAGAGCATGCCGGGCCTCCCTGGCACGGCCCTCTGGGCTCTGT	6737
gGluArgAspArgAspArgGluArgAspArgAs	180
6738GGAG	674
1809 luargGluLys-SerIleLeuThrSerThrThrThrValGluHisAlaProlleTrpArg	parg 1828 GTCC 6798
1829 ProGlyThrGluGlnSerSerGlySerSerGlySerSerGlyGlyGlyGlyGlyGlyGlyFerSer	184
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849	1865 ATGC 6879
6820 CGICCACCACCACCACCACCACCACCACCACCACCACCACCA	
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1883 MetLysGlyIleIleThrAlaValGluProSerLysProThrValLeuArg	
6931	STGGG 6969

411 AsnMetAsnGlyLeuMetA ::: 2362 TCCAGGCGCAGCAGCATG	4 0
431 MetTrpSerGluGlnGlul ::: 2419 TCCCACAGTGAGAAACC	MetTrpSerGluGluGluLysGluThrPheArgGluLysPheMetGlnHisProLysAsn 450
451 PheGlyLeuIleAlaSerl 2479 ACCGCCCCCTGTGCCT	PheGlyLeuIleAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyr 477
471	TyrLeuThrLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArg 49
2538 CAGCACCCCCACCACC 491 GlyLysSerGlnGlnGln(
510	CGAAGCCCTGAGCCGCAGCAGTCACGTGFTFTACCTCCCCACCCCCGGATGCTGAAGGCCGAA SSS
	rSerGlyGluAspAsnAspGluLysGluAlaValAlaSerLy 56
567 sGlyArgLysThrAlaAs 	erGlnGlyArgArgLysGl AAAGG
582 eThrArgSerMetAlaAs 	eThrargSerMetalaasnGlualaasnSerGluGlualaileThrProGlnGlnSeral 602
602 aGluLeuAlaSerMetGl :: 2889 AGAGCTTCAGATCGCAAA	aGluLeuAlaSerMetGluLeuAsnGluSerSerArgTrpThrGluGluGluMetGluTh 62: AGAGCTTCAGATCGCAAGCCCATCTCTGCAGGCACCCCACATCTCCAGAA 29
622 ralaLysLysGlyLeuLeuGluHisGl. 	uGluHisGlyArgAsnTrpSerAlaIleAlaArgMetValGl 64 TGAGCATGAGCCGTGGTCCCAA29
642 y	nCysLysAsnPheTyrPheAsnTyrLysLysArgGlnAsnLe 66
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682 sLysLysLysAlaProAl :::::: 3030 GAGGAGAAAGAGAGAGA	BLYSLYSLYSALAProAlaAlaAlaASerGluGluAlaAlaPheProProValValGluAB 7
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The invention relates to modulating inflammation or apoptosis in an immune cell. The method involves contacting the cell with a compound that modulates RKC (KappaB binding and puterive recognition compounent of the compound to the modulative and the cell with a compound that modulate and the cell is modulated. Also provided are methods for modulating immune cell is modulated. Also provided are methods for modulating immune cell is conflicted on a activation and for inhibiting metastatic growth of a tumour cell. The method is useful in modulating an immune response, such as inflammation or apoptosis, using agents that modulate KRC activity. The method may also be used in identifying agents that modulate KRC activity which can be used for treating or preventing disorders associated with an aberrant cell proliferation or survival, such as a sociated with an aberrant cell proliferation or survival, such as a sociated with an aberrant cell proliferation or survival, such as a second sorders or graft-versues-host disease. The present sequence represents a cDNA encoding a human KRC protein, a DNA binding protein. KRC is a member of zinc finger proteins that bind to the KappaB motif
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		GGCTCGGACTCGTCGGCGCGCGCGCGCGCACGCGGCGGCCTGGTCGGCGTCACGACG	1795 GluArgGluArgABD 1/99 1795 GluArgGluArg		TGCT				1843 GlyGlyGlyGlySerSerArgProAlaSerHisSerHisAlaHisGlnHisSerPro 1862		1863 IleSerProArgThrGlnAspAlaLeuGlnGlnArgProSerValLeuHishsn 1880	GICGGICCGCIIGICCGGCIICGGCCIICGCCCCCCCCC	1881 ThrdlyMetLyBG1y11elleInralavaloudrideLyBf10111		14236 TCGGCCACCGGGACACCGGCCGGCCGCCCCGCTCCCAGGTGCGCATCGCC 14177							1947 gProGluArgProArgAlaAspThrdlyHisAlaPheLeuMlaLyBFroFroAlaArgse 1997			1985 ProProVal	cerecaceaec			2012 ProAlaProProAlaSerAlaSerAspProHisArgGluLysThrGlnSerLysPr 2030	ACGCCCGAT-GCGCCCCCTTCTCGACCGCGGCCAGCCAGGGCCGGTCGTCGCCCCGGACAGGCCCCGACAGGCCCCGGACAGGGCCGGACAGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGTCGGGGTCGGGGTCGGGGTCGGGTCGGGGTCGGGTCGGGGTCGGGGTCGGGGTCGGGGGG	::: :::
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703	CAGCCGCTCGGCTCCCGGTCGGCGAGC
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2070 1	uProLysHisLeuGluGluLeuAsp-LysSerHisLeuGluGlyGluLeu 2086 UProLysHisLeuGluGluLeuAsp-LysSerHisLeuGluGlyGluLeu 2086 UProLysHisLeuGluGluLeuAsp-LysSerHisLeuGluGlyGluLeu 2086 UProLysHisLeuGluGluLeuAsp-LysSerHisLeuGluGluGluCluLeu 2086 UProLysHisLeuGluGluLeuAsp-LysSerHisLeuGluGluGluGluCluLeu 2086 UProLysHisLeuGluGluLeuAsp-LysSerHisLeuGluGluGluGluCluCluCluCluCluCluCluCluCluCluCluCluCl
087	hrgproLysGlnProGlyProValLysLeuGlyGlyGluAlaAlaHisLeuP 2104
2104	roHisLeuArgProLeuProGluS 2112
2112	erGlnProSerSerProLeu
212	GCCGGGCGAGGTCGTCGCGGGGAGGACGCCCTGGGTCAGGTCAGCCCCCCCC
2131	rgvalvalthrLeualaginHisileSerGluvalilethrGlnAspTyrThrA 2149
149	rgHisHisPro
2159	LeuProAlaProLeuTyrSerPhe-ProGlyAlaSer
2172	
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2271	2274 PhepheserL 2274
12744	CGACCGCCTCCTCGACGGCCTGGACCACGATGCGGGCTGACCTCGATGAACGCGT 1268
12684	4 CGTAICCCGACTCCGCCAGCCGCGTCACGGCGTCGGCGAAGCGCACCGTCTCGCGCAGGI 12625

1305 AGCCCCGCCGCGCGCGCGCCGCCGCCGCCGCCGCCGCCGC	
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18608 AAGAGGGGGTCCGGAACCGCAGCAGC	GAGCTGCGGGTTGTCCGGAACCGCAGCAACATCCCAGGTTCCGCGTCGGTGTCTCCGTCGTTCTCTCGTGTTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCTCT	- -
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589GluAlaAsnSerGlu		qa ,
18548 CACGAACCGGTCCGGCGACTCGGCCTGCGCACAGCGGATCACGCCCCACATCGCGGCGGC		λō
603 uLeuAlaSerMetGluLeuAsnGluSerSerArgTrpThrGluGluGluHtrAl		qa
18488 GCCGGGTTCGGGAACGCTGTCGTCGACGGTGACCGCCCGGACGTGGCGATGAC	ACGGTGÁCCGCGCCGGACGTGGCGATGAC 18429	۵
623 aLysLysGlyLeuLeuGlu		<u>ස</u>
18428 CAGCGTCGCGCCCGGCAGCTCCTCTTTCGAGCCAGGGGCGCACGAGCGCGGCGGCCCA	AGCCAGGGGCACGAGGGGGGGGCCA 18369	λõ
639 gMetValGlySerLysThrValSerGlnCysLysAsBsnPheTyrPheAsnTyrLysLysAr		qū
18368 GAGGACGCCGTGGCGGGCCGCGGGGGGGTCGTCGCCTTCGGGTTCGTAGCGGACCAG	TCGTCGCCTTCGGGTTCGTAGCGGACCAG 18309	ò
659 gGlnAsnLeuAspGluIleLeuGlnGlnHisLysLeuLysMetGluLysGluArgAsnAl	LysLeuLysMetGluLysGluArgAsnAl 679	q
18308 ARCCGCGTCGGGTTCGGGTTCACCGGACCGCAGGACGGCGTCGAGGTCCGAGGCATCGGC	AGGACGCCTCCAGGTCCCAGGCATCGGC 18249	ò
679 aArgArgLysLysLysLysBAlaProAlaAlaAlaSerGluGluAlaAla	AlaSerGluGluAlaAla 695 	ପ୍ର
18248 CGCGACCACAGGGCGGTCGAGCGGGCTGGGTAGCGGTCCCCAGTCCAGAGCGTGCAG	GCTAGCCGTCCCAGAGCGTGCAG 18189	ò
969	701	ପ୍ଧ
18188 CICGCCCTCGCCACCGCGCGCTTCGGGCTGCTCCCACTTCTCCCCCGGTGCTGCGGACGAC		70
702 -AspGluGluMetGluAla	707	qa
18128 CAGCGAGTCGACCGTGGCGACCAGGTGGCCTGCCGGGTCGGTGACCCGCAGGGCCATCGC	IGCCGGGTCGGTGACCCGCAGGCCATCGC 18069	٥٥
708SerGlyValSerGlyA8nGluGluGluMetValGluGluAl		qq
18068 GTCCGCCCGGCGGCGCGTCGCCACCGCACCGAAGTCGCCCCGGAAGGCGTGCAGGGT	zaccdadgrcgcccgdaddcgrgcagggr 18009	8
722 -GlualaLeuHisalaSerGlyAsnGluValProArgGlyGluCysSerGlyPro	739	DP DP
18008 cacegrerrccacededacedeacrrrcccccecceecrcecceaegececcaeger		8
740AlaThrValAsnAsnSerSe	ProHisTh	qa
17948 GAGCGTCTGCGCCACGGCGTCGAGCACCGGGTGGAACGCGTAACCTTCC	TC	ò
756 rGluAlaAlaLy8AspThrGly		<u>අ</u>
17894 GrccGccGcGArGGACACCrcGGCGTAGAC	: Greegegegeargandactegegeraganegregreerregegegegegegegerag 17835	ò
764GlnAsnGlyProLysProProAlaThrLeuGl		qa
17834 GGCCTGGAACGACGGTCCGTACTCGTA-ĠĊ	GGCCTGGAACGACGGTCCGTACTCGTA-GCCCTGCTCGAGGCCGTCGTAGTGGTCGT 17776	δ
775AlaAspGlyProProProGlyPro-	yPro	qa
17775 cgagcggardcggacdgcgr-ccrccgc	cgagcgggargcggacggcgr-ccrccgcaccacrgcrccgrarcccrcggacccgcg 17717	λο
186	786	qq
17716 GCGACGCCCTGCGCGAGCGTCCCGGTGGC	GCGACGCCCTGCGCGAGCGTCCCGGTGGCGTGCGACCACTGCGCGTCGGCGAGGTCG 17657	ò
787ArgArgTl	-ArgArgThrSerArgAlaProlleGluProThrProAl 799	ପ୍ର
17656 GCCACGTCTTCGGCGGCGTGGACGTCGAT	GCCACGICTICGGCGGGGGGACGTCGATCGTCCGCCGCCCCCCACTCGTCGGGCGCGCGC	ò
799 aSerGluAlaThrGlyAlaProThrProF		අ
17596 ACCGACATGCGCAGCAGCGCCCCCCCCC		<i>\delta</i>
	cl.,.rhralaalaalaproProValGluGl 839	

36 TCCTCCAGGACCG	Η α
CCCACGACGTGT 17	17
859 uGluProValLysSerGluCysThrGluGluAlaGluGluGl 87	1 873 17381
73 YProAlaLysGlyLysAspAlaGluAlaAlaGluAlaThrAlaGluGlyAlaLeuLysAl 8	
893 aGluLysLysGluGlyGlyBerGlyArgAlaThrThrAlaLysSerSerGlyAla	- 911 C 17270
912	- 915 G 17210
930 uGlydspLysAsnArg	-
942 oserLeuLeuThrProThrGlyAspProArgAlaAsnAlaSerProGlnLySPr 9 	
960 oleuAspLeuLysGlnLeuLysGlnArgAlaAlaAlaIleProProIleGlnValThrLy 9	> F
9 .HHi 9	982
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982 sGluProProArgGluAspAlaAlaProThrLy8	
996 oProAlaproProProGlnAsnLeuGlnProGl 1	
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1044 YASPPROPROCYSTRPThrSerGlyLeuProPheProVa ::	Va 1057 .TG 16643
1057 lProbrogGluVallleLysAlaSerProHisAlaProAspProSerAlaPheSerTy 1057 lProArgGluVallleLysAlaSerProHisAlaProAspProSerAlaPheSerTy 11	Ty 1077 16597
1077 rAlaProProGlyHisProLeuProLeuGlyLeuHisAspThrAlaArgProValLeuPr 	
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18668 CAGCGCGGAAGGTGCGAGCGCGAGGCGCG	qa	207 ALGABILITERIANG GASTILITERIA SOCIETIAN SOC	3 €
571ThrAlaAsnSerGlnGlyArgArgLysGl	ò		a è
18728 GGCCACGGAGTCGATCGCCCCGTCACCGGCAC	2 A	267 HisGluAsnIleLys1leAsnGlnAlaMetArgLysLysLeulleLeuTyrPheLysArg	8 8
559 pGluLvsGluAlaValAlaSerLvsGlvA	8 8	15	q _Q
542LysGluAspLeuLeuLysGluLysThr	& A		ò
18848 CTTCTGCGGATACATGCCGAGTGCGAGGAGGA	ପ୍ର	19766 GTCGGAACCCGGCGCGCGCGCTGTGACCAGCAGCAGCAGCACCCGTG	; A
524 GluLysGluLysGluAlaGluLysGluGluGlu	ò	13080 GLMAGCCGCGA1CICCGCGGACGCCCCAAGICCICGAICICCCGCGCAGCIC	3 2
18908 GGCGTCCACGTCCGGTCCGACCGCCGTGACGAC	අු		හි සි
517GlnGluGluLysAgpGlu	ò	19880 GGTCAGCGGGCCCGGCCCCTCCAGCGCCGCGTCGCGGGCGG	aa (
18968 GACCGCGATCGGCGCGAAGGCTCCCTGGAACACAC	ą q	204 AlaAlaLysProProGluProGluLysProValSerProProDile ::: ::: :::	ð :
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	ପ୍ର	19991 GTGCAGGTTCCACGCGCGTCGACCTTGGCGCGCAGCACCTGCTCCACCGCCGGCTCGTC	ч
476 AsnGluAsnTyrLysSerLeuValArgArgSer	ò	173 []eGlnasnMer Asnardvallan	
19142 GACGGCCGCCATGCCGACACCGCCTGCCGCTGC	qq	Qy 169LysGluGluLeu 172 Ander Charles C	~ L
461bysThrValAlaGluÇYę	ò	20111 AGCCAGCGCGTTGAGCGACTCGTTCGCGGCCGCGTACACGCCCTGCCCCGGGCCGGCTAG	~
19184 GGGGTGGCAACACTCCGCCCCGCTCGCC	à a	157 AspèroGluLeuGluLe	Ġ
	<u>a</u> :	Db 20171 CTGCGCCCACCCCATCCGAGCGCCTTCGCGGGAAGGCCGCGCGCG	ы
423 TyrLysAspArgGlnValMetAsnMetTrpSer	<i>&</i> 1		ò
19259ACCGGTCTC	d d		3 43
405 GlnArglleLysPhelleAsnMetAsnGlyLev	<i>λ</i> ο	20264 GCTGTCGAACAGTGCGAGCGCCCGGTCGGCGCGCGGCGACCCCCGGTCCGGGCGAT	a 6
Ö	3 8	109 ArgleuGluLeuLeuProAspProLeuLeuArgProSerProLeuLeu	ò
19370 GCCGICCICGGGGGGGGGGGGGGGGGGGGGGGGGGGGG	<u> </u>	Db 20303GGACAGCGGGAACACGACCTCACCGCCGCGCGCAGGGC 20265	
	ð i		
19430 GTATCGGCCCGGAAGICCCCGGCGICCCGGCAC	qa	Db 20342 CGGCACGAACTCGGCCCTGCGCAACGCTGAT	, H
361 Leuser	ò	20402 IGCCGCCTGCCCGGCGGCGCGCGCAGCTTCGCCCTGACCATGCCGCGCAGGACCTC	
	3 8	49	ò
19550 TCCCAGCTCCCAGCCGGTACCGGGAGCCGGT	8 8	Db 20454 CGACCTGGTCGGACTCGGACCGGCGAGCCGGTCGACCTCGACCT 20403	1
327 SerLysValArgGluTyrTyrGluLys	<i>à</i> 1	32ArgThrHisThrAspValGlyLeuLeuGluTyrGlnHisHisSerArgAsp	ò
19592GTGCCTGCCCGGCT	අධ	OY 13 AFGALELINICALUFIOAGGIYETOFIOAISSELEGUSETIYETOVALOLILIGALA 31	දී සි
307 GluAlaLeuGluLysLysValGluArgIleGlu	δ		_

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λ:	343	GlnArgGluLeuGlnGluArgMetGlnSerArgValGlyGlnArgGlySerGly 360
q	19490 GA	ACCTCGCGCAGGATCTCCCCGAGCCGGTCGTCACCCGCCTCGCCGAGGTCGAACGGGGC 19431
*	361 Le	euser
ą	19430 GT	GIATCGGCCCCGGAAGTCCCCGGCGTCCCGCAGGTCTTGCCCATCTCGACGAAGAC 19371
~	370 Hi	HisGluValSerGluIleIleAspGlyLeuSerGluGlnGluAsnLeuGlu 386
ą	19370 GC	ccarcercaccadeadrecacadacrearcadacadreceargadeagreada 19311
⋩	387 Ly	LysGlnMetArgGlnLeuAlaVallleProProMetLeuTyraspAlaAspGln 404
ą	19310 CA	accacgrececececedecrecegrecerecedaaacrrecegeceaa 19260
≿	405 G1	GlnArgileLysPheileAsnMetAsnGlyLeuMetAlaAspProMetLysVal 422
ą	19259	
*	423 Ty	TyrLysAspArgGlnValMetAsnMetTrpSerGluGlnGluLysGluThrPheArgGlu 442
q	19223 GA	AGACCGAGCGC
≿	443 Ly	LysPheMetGlnHisProLysAsnPheGlyLeulleAlaSerPheLeuGluArg 460
ą	19184 GC	3CGGTGGCCAACACCTCCGCCCCGCTCGGCGGGCCAGCGC 19143
*	461	LysThrValAlaGluCysValLeuTyrTyrTyrLeuThrLysLys 475
ą	19142 GA	Acegececcarecearaccecrececreceredeareacaccaacteaccece 19083
~	476 AE	snGluAsnTyrLysSerLeuValArgArgSerTyrArgArg489
ą	19082 CA	CAGCCCCGCGAGATCGTGCAGCGCTAATGCGCCGTGGTGTAGGCGATGGGCAC 19029
⋩	064	496
ą	19028 GG	SCGCGCGCGTCGCTCCAGCCGTCCGCGCACTCGTGCGAGGGCCGGTGATCGGT 18969
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~	524 Gl	luLysGluLysGluAlaGluLysGluGluLysProGluValGluAsnAsp 541
ą	18848 CT	CTTCTGCGGATACATGCCGAGTGCGAGGAGGACGTCGCGGAAGTTGACTCCGGTGGCGCG 18789
~	542	LysGluAspLeuLeuLysGluLysThr-AspAspThrSerGlyGluAspAsnAs 559
ą	18788 CP	CACGTCCACCCGGACCTCGCCCGGAGCGGCTGCTCGACGTCGGGGCGGGC
⋩	559 pC	pGluLysGluAlaValAlaSerbysGlyArgLys570
<u>۾</u> :	18728 GC	SCCACGGAGTCGATCGCCCCGTCACCCGGCACGTTGGGTGCCTGCC
≿	571	ThralaAsnSerGlnGlyArgArgLysGlyArgIleThrArgSerMetAlaAsn 588
ą	18668 CA	CAGCGCGGAAGGTGCGAGCGCGGGAGAGGCGCGCGAAGACGTCGTCGTCGCGCGAAGGC 18609
<i>\</i>	588	885

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           20561 GricchicroccogriccogriccegaacGricacegraaggraaggraaggraace-----Carac 20508
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                1113 GGCCACCGATCCGGCCGGCACGGCCGCGAGTCCGCGACCGCGAACCTGGGCGAA 21354
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                                                                                                                                                                                          2271 PhePheSerLysLeuThrGluSerAsnSerAlaMetValLysSerLysLysGlnGluIle 2290
                                             hrSerValLeuGlyGlyGlyGluAspGlyIleGluProValSerProProGluGlyMetT 2232
 -- ArgSerProGluProAsnLysT 2212
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                                                                                             2232 hrGluproGlyHisSerArgSerAlaValTyrProLeuLeuTyrArgAspGlyGluGlnT
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                                                                                                                                                                                                                                                        2366 AsnAlaPheAsnProLeuAsnAlaSerAlaSerLeuProAlaAlaMetPro-----
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                                                                                                                                             2252 hrGluProSerArg-MetGlySerLysSerProGlyAsnThrSerGlnPro---ProAla
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                                                                                                                      21302 GAACGĊĊĊAGTCGGÄĠŤ------
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This sequence represents a fragment of the Saccarapolyspora erythraea genome, designated eryA. The polypeptides encoded by this region are involved in the biosynthesis of the polyketide segment of erythromycin. eryA is organised in modules and each module takes care of one condensation step. The precise succession of elongation steps is dictated by the genetic order of the modules. This fragment may be specifically altered such than novel polyketide molecules of desired structure are produced. Three types of alteration may be produced; those inactivating a single function in a module which does not arrest acyl chain growth; those inactivating a single function in a module which does not arrest acyl chain growth; and those affecting an entire module. The mutations may be introduced by gene replacement. (Updated on 25-MAR-2003 to correct PN field.) (Updated on 27-AUG-2003 to correct PN
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  osynthesis of specific polyketide analogues esp. erythromycin cpds. introducing altered biosynthetic gene-contg. DNA into microorganisms.
                                                                                                                                                                       Saccarapolyspora erythraea; eryA; biosynthesis; polyketide; module; erythromycin; condensation; elongation; acyl chain growth;
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P-PSDB; AAR44430, AAR44431, AAR44432.
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3. .10199
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1417 GTTCGGTGAGCCCTTCGCGGGCTCCTCCCCCCCC	1960 LeuAlaLyPeProProAlaArgSerGlyJeuGluProAlaSerSerProSerLyeGlySer 1979 22433	21533 GACGCACGCAGCAGATGCCCAGATAGGCGACGCCCGCCCG
1895 eTICHERANGE CETTCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCC	CGAT 23379 snTh 1699 GCAC 23319 erSe 1719 TCGA 23277 roHi 1739 CGCA 23226 etAs 1756 GG 23170 GG 23170 GG 23170 1781 CGCC 23050 1781 CGCC 23050 1781 CGCC 23110 1810 GCGT 22871 hrVa 1821 CAAC 22811 1830 1830 1843 TGAC 22631 Hiss 1861 GCGG 22631 GCGG 22631 GGCG 22631	22485 aphe 1959 c 22434
	1685 ElleABROGCTTCCCGGCG	ualaProargValalaArgProGluargProargAlaAspThrGluargProargAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargProArgAlaAspThrGluargPr

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24254 GGGCCCCGGCC 24243
1372 rgargGlualaLysLeuLeuLysArgGluGlyThrProProProProPro-SerArg 1391 24242 GCCGCATTGGCCGCACCGATAGCCGACCCAAATCCGCCGCAGCCGC 24184
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1456 ProLeuLysTyrAspThrGlyAlaSerThrThrGlySerLysLysHisAspValArg 1474
1475 SerLeulleGlySerProGlyArgThrPheProProValHisProLeuAspValMetAla 1494 23914
495 AspAlaArgAlaLeuGl :: 896CA
1515 AlaSerSerGlyGlyGtySerIleAlaArgGlyAlaProVallleValProGluLeuGly 1534
1535 LysproArgGlnSerProLeuThrTyrGluAspHisGlyAlaProPheAlaGlyHisLeu 1554 :::
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1616 gGlyValSerGlyValAsp
1626 rHisIleProLeualaPheAspProThrSerIleProArgGlyIleProLeuAspAl 1645
1645 aAlaAlaAlaTyrTyrLeuProArgHisLeuAlaProAsnProThrTyrProHisLeuTy 1665 23477 TGAAGCTCGCTTTGCAGCGCCGCGGGCTCGCCGGGGAATCCAGGGCCACCGCTGCCA 23418
1665 rProProTyrLeulleArgGlyTyrProAspThrAlaAlaLeuGluAsnArgGlnThr11 1695

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------AlaAlaGlyProArgGlyIleI 1732 |||:::||| ACACGCAGGCCGTTCGGGTGTCGCCGGTGTCA 31100 SGTGGCGGCGTGGCCGTGGCCGGTGTCCGC 31380 CCTGGCCGTCGTGCGCGCAGCGCCATCAACC 30830 SCTCCGCCGGGCGGCGTCCCGCGTTCGGCGT 31272 scacerceregagesecceesecea 31320 LysLeuThrSerThrProArgGlulleAlaLys 1594 3GGGCGAGGGCGTCGCGTACTCCTCCTCGAAC 30764 30789 |||||||| TCCGCCAGGCCCTGGCCGACGCGCGCTCACGT 30920 Aceccaces and a social contra social de la contra c rgcaccrccrcaagggccrgcggaacggcg 30596 |||||| 3GGTGCTG-----CCGAAGACCCTGCACGTGG 31154 BluHisHisProHisProIleSer-ProTyrGl 1612 AspleuTyrArgSerHisIleProLeuAlaPh 1632 ProLeuAspAlaAlaAlaAlaAlaTyrTyrLeuPr 1652 yrprodisleuTyrProProTyrLeuIleArgG 1672 SnArgGlnThrileIleAsnAspTyrileThrS 1692 isAsnThrAlaThr----- 1701 euProThr ------AlaProGlnPro 1767 euSerProglyGlyProThrHisLeuThrLys 1787 3luProThrProArgLeuGlnGluGlySerLeu 1574 covalLeuValProPro-ThrProGlyThrPro 1751 3luArg---- 1798 LysSerIleLeuThrSerThrThrThrValG1 1822 crcccckc-----

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5 A	27439 GGCCGAGGTCCTCGCCGGAGCTCCGCACGTCCGCGTCTTCTCTCCACCCT 27498	8 8	1030AlaAs
ò	800 erGluAlaThrGlyAlaPro	7 음	:: 28537 GGCCCACCACGGAGCCGAA
q	27499 CGAAGGCACCTGGATCACCGAGCCGGTGCTCGACGGCACCTACTGGTACCGCAACCTGCG 27558	λō	1045 spProProCysTrpThrSe
ζ		q	:: 28597 AGCCACCC
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ò	SerproSerAlaProProProValValProLysGluGluLysGluGluGLuTnrAlaA	qq	28645 carcececereceacere
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ò	laAlaProProValGluGluGluGluGluGluGluGluGluGluGluGluGlu	eg C	28705 CG
q		ò	1105 roProProLeulleSerSe
ò		qa	28735 CACCGCCGCCCACCGGG
qu	27739 CIGGGCCAACGCCCICACCAICGACTGGGCGCCCAICCTCCCCACCGCAACGGGCCACCA 27798	λö	1116 erValLeuGluArgGlnI
ò		qq	28795 CATCCTGGGCGCGAAGAC
qq	TGGCTGCAGAGCTCCGCGCC	ζ,	1136 roTyrSerGluHisAlaL
ò	877	qu	28855 GCTGGACGCCTTCGTCCT
QD	27859 CACCAGCGCCGCCGACGACTGGCGTTACCGCGTCGAGTGGAAGCCGCTGACGGCCTCCGG 27918	Š	1148 alThrMetGlyLeuProL
λō		q ₂	::: 28915 CGTCTACGCGGCGGCCAA
qq	27919 CCAGGCGGACCTGTCCGGGC	δ	1165 erGlyValLysGlnGluG
٥'n	006 006	qa	28971GGGGCGAGACGGCGAC
qq		λō	1184 euGlyValProThrAlaG
ò	901GlyArgAlaThrThrAlaLysSerSerGlyAlaProGln- 913	qa	29026 CCGGGGCGCCGACGACGC
qq		Š	1196
ò		q	29086 CGCCCTGGACGAACTGGC
q		λ	1196 rgGlyThrAlaLeuGlyS
ò		qa	29146 TGTCGACTGGGAGCGGTT
qq	CGGCGACGCCGGAATCAAGGCGCCCCTGTGGTCCGTCACCCA	ò	1211GlylleProSerThrA
ò	LeuThrProThrGlyAspP 	qa	29205 ACGCCGTCCCGGAGGCCC
qq	28195 dececedetriciarcearcererearacecideceacecedacegecear 28248	λō	1220 spSerAlalleThrTyrA
ò		qu	29265 ACGCCGCCGTGGCGCCG
q		ζ	1231HisGlyThrProf
ò	<pre>fhrLysValHisGluProProArgGluAspAlaAlaP</pre>	qq	29325 AGCCCGAGCGCCGGCCGC
qq		ð	1248 lyGluAspSerProSerA
ò	991 roThrLysProAlaProProAla	ą	29385 GCCATTCCTCCCCCGACC
qq	28357 CGCACTCTCCGGCGCCACCGGCGAGGACCAGATCGCCATCCGCACCGGACTCCACGC 28416	ð	1268 alileTyrGluGlyLysi
ò	1006GinProGluSerAspAlaProG 1013		

||||| ::: ||| ::: ||| 39384 ::: GGTACTGGCAGCGTCGCGGCATCCGATGAGCCCCGACCG 29085 CCAAGGCCCTGAGCCACGACGACCTTCGTCGCCGTGGCCGA 29145 |||| || || ||| rrdeg-cccgcgrrcacgargrcccarccagccrrcrgcrcg 29204 cedecadecectresces acces as a second con contraction of the contractio |||| gaccgggcagtcgtcgccgctgccgcgalcaccgcgctccccg 29324 SLYSGlyHisValLeuSerTyrGluGlyGlyMetSerValThrG 1288 CAAGCGGCGAGGTCCTCGACGACCTGCTCCGCGGCACTCC 28854 TCTACTCCTCGAACGCCGGGGTCTGGGGCAGCGGCAGCCAGGG 28914 SerValProGlyGlySerIleThrLys------ 1210 -----ValProSerA 1220 Argdly----- 1230 AlaAspVal---LeuTyrLysGlyThrIleThrArgIleIleG 1248 rArgLeuAspArgGlyArgGluAspSerLeuProLysGlyHisV 1268 GCGGCGATCCGCTGGACGTCACCGGCCCGGAGGACATCGCCCG 28794 AACACCTCCTCCTCGTGAGCGGGGAACAAGCCCCCGG 28596 GlnLeuSerProArgGlyGlnAlaGlyProProGluSer---L 1184 GlnGluAlaseryalLeu-----1195 cescecaccesascectesses de casececas de 18536. ---AACTCACCGCCGAACTCACCGCATCGGGCGCCCCGCGTCAC 28644 ThralaargprovalleuproargproproThrIleSerasnP 1105 SerAla-----LysHisProS 1116 IleGlyAlaileSerGlnGlyMetSerValGlnLeuHisValP 1136 LysAlaproValGlyPro------ 1148 LeuPro------MetAspProLysLysLeuAlaProPhes 1165 splysGluAlaPheAlaAlaGluAlaGluLysLeuProGlyA 1045 serGlyLeuproPheProValproProArgGluValIleLysA 1065 AspproSerAlaPheSerTyrAlaProProGlyHisProLeuP 1085 roArgGlyLysSerArgSerProAlaProPro----- 1029

8	25462 CTTCGGGATCTCGCCGCGAGGCCCTCGCCATGGACCCGCAGCAGCGACTGTCCCTCAC 25521	526]"
ò	209	22720
QQ	25522 CACCGCGTGGGAGGCGATCGAGAGCGCGGGCATCGACCCGACGGCCCTGAAGGGCAGCGG 25581	00407
È	218 ProlleGluSerLysHisArgSerLeuValGlnIleIleTyrAspGluAsnArgLysLys 237	529
QQ	:::	26
ò	238 AlaGluAlaGly 249	Qy 539
q	AGCTGGAGGCCCACCTGG	26586 GGCCGGTGTCCGC
à	250 ProGlnValGluLeuProLeuTyrAsnGlnProSerAspThrArgGlnTyrHisGluAsn 269	Qy 540AsnAspLysGluAspLeuLysGluI :: :: ::
q	GACCGGCCCTGACCGTGGACACGGCCTGCTC	Db 26646 ACGCGGACGTCGTACGGACGTGGATCCGGCGGTGGCCGCCC
ò	IleLysIleAsnGlnAlaMetArqLysLysLeuIleLeuTyrPheLysArqArqAsnHis	Oy 555 lyGluAspAsnAspGluLysGluAlaValAlaSerLysGlyA
d d	: TGCAGGCCCTCCGCAAGGGCGAGTGCGACAT	Db 26706 GTACGGCGATGGAGCACCGCGCGGTCGCGGTCGGCGACAGCC
è		Oy 575 lnGlyArgArgLysGlyArgIleThrArgSerMetAlaAsnG
7 A		Db 26766 cccrecedareccesaagaacregracecacarcerce
ઠે	ArgArgAlaLysGluSerLysVal	Qy 595 laIleThrProGlnGlnSerAlaGluLeuAlaSerMet
අු	25858GGACCTGTTCGTGCAGTTCAGCCGGCAGCGGGGCTGGCCGCGGACGGG 25905	Db 26826 TCGTCTTCCCCGGCCAGGCACGCAGTGGGCCGGCATGGGCC
Ś	AraGluTvrTvrGluLvsGlnPheProGluIleAraivsGlnAraGluLeuGlnGluAra	Oy 613 erArgTrpThrGluGluGluMetGluThrAlaL
7 8		Db 26886 CACCGGAGTTCGCTGCCTCGATGGCCGAATGCGAGACCGCG
i è		Cy 630 isGlyArgAsnTrpSerAlalleAlaArgMetValGlySerI
3 8		Db 26934 GCTACGTCGACTGGTCT
3 8		Qy 650 ysAsnPheTyrPheAsnTyrLysLysArgGlnAsnLeuAspC
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9	CAALGGALCTCGCGGTCGTCGCGGCAGCGCGGTCAACCAGGACGCCGCCAG	Qy 670 ysLeuLysMetGluLysGluArgAsnAlaArgArgLysLysL
à ·	rgGlnLeuAlaVallleProProMetLeuTy	Db 26972GAACC-CGGCGCACCCACGCTCGACGCTC
8	26059 CAACGGCCTCACGGCTCCGCACGGGCCCTCCCAGCGCGCGTCATCCGACGGGCCCTGGC 26118	Qy 690 laSerGluGluAlaAlaPheProProValValGluAspGluG
ò	400 rAspAlaAspGlnGlnArgIleLysPheIleAsnMetAsnGlyLe 415	27016
q	26119 GAACGCCCGGCTCGCGCCGGGTGACGTCGACGTCGACGCCGCACGGCACGGCCACGCG 26178	710 alserglvasnelnelnelnestvalelnelnalae
ò	415 uMetAlaAspProMetLysValTyrLysAsp-ArgGlnValM 429	27038
qa	26179 GCTCGGCGACCCGATCGAGGCGCAGGCCCTCATCGCCACCTACGGCCAGGAAGAGAGCAG 26238	730 luValPro
ò	429 etAsnMetTrpSerGluGlnGluLysGluThrPheArgGluLysPheMetG 446	575757757#454575554557#7477577#5 98076
q	26239 CGAACAGCCGCTGAGGCTGGGCGTTGAAGTCGAACATCGGGCACAC 26286	741
ò	446 lnHisProLysAsnPheGlyLeuIleAlaSerPheLeuGluArgLysThrValAlaGluC 466	77145
ą	26287 GCAGGCCGCGCGCGCTGTCGCAGGTGCTCCAGGCCGATGCCCACGCCACGGACT 26346	C\$T/7
ò	466 ysValLeuTyrTyrLeuThrLysLysAsnGluAsnTyrLysSerLeuValArgArgS 486	70 YB:::: : : : : : : : : : : : : : :
Op	26347 GCTGC 26351	607/7
ò	486 erTyrArgArgArgGlyLysSerGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnG 506.	Db 22255 CGACGGACTCTCGTCGTCGAAGGGCCCCAC
qq	26352CGAAGACGCTGCACGTCGACGAGCCCTCGGACCAGATCGACTGGTCGGCGGCAC 26406	783 roThrProProArgArgThrSerArgAl
δ	506 InGInGInGInProMetProArgSerSerGInGluGluGluLysAspGluLysGluLysG 526	23
QQ	26407 GGTGGAACTCCTCACCGAGGCCGTCGACTGGCCG-GAGAAGCAGGACGGGGGCTGCGCC 26465	

GCACGTCGTCCTGGAGGAGG 26525 ccccccrccrccacaccc 26705 ||||::: CCGGGAGGCACTGCGGGACG 26765 ----GluLeuAsnGluSerS 613 |||||| CGCCGAACTCCTTGACAGCT 26885 AGCCGTCGTCCGACAG---- 26971 SLysLysAlaProAlaAlaA 690 ::::::: TCGACGTCCAGCC--- 27015 GCATCACCCCCAGGCCGTC 27085 ||||||| ACCGCCACCGTCGTCTCCGG 27318 CGGTGGCGGTGTGCCGT 26585 SATCGGGCAGCTCGCCGCGT 26645 SGACGTGGGCCGGGTGGCGT 26825 :::::: TGACCTTCGCTGTCATGGTC 27037 yGluCysSerGlyProAlaT 741 |||:::|||:::| CCGGTGCACTCA-CCCTCGA 27144 GCGCCCACCTCGCCGGCAA 27204 CTGAAGCGACTGAGCGACTT 27264 pGluIleLeuGlnGinHisL 670 aLysLysGlyLeuLeuGluH 630 539 JLYSThraspaspThraerG 555 yargLysThrAlaAsnSerG 575 nGluAlaAsnSerGluGluA 595 rLysThrValSerGlnCysL 650 uGluMetGluAlaSerGlyV 710 aLeuHisAlaSerGlyAsnG 730 rProHisThrGluAlaAlaL 760 -----ABDThrG 763 pGlyProProProGlyProP 783

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                                                                                            33391 CACGGGTACGGGACGGCACGGCCTCCTCCCGGCCGATCTCGACGCGCTCGACGC 33450
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                                                                                                                                                                                                                2450 rpGluAspArgProSerSerAlaGlySerThrProPheProTyrAsnProLeulleMetA 2470
                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Streptomyces venezuelae; Sv; narbonolide polyketide synthase; PKS; narbonolide PKS; narbomycin biosynthesis; picromycin biosynthesis; procromycin biosynthesis; procromycin biosynthesis; prkS gene cluster; picromycin, glycosylation; hydroxylase; PicK; deoseamine biosynthesis; desosaminyl transferase enzyme; antibiotic; narbonolide synthase; gene;
                                                                                                                                   rgLeuGlnAlaGlyValMetAlaSerProProProGlyLeu-ProAlaGlySerGly
                                                                                                                                                                   -----AspCysAsnArgArg-----ThrProLeuThrAsnArgValT
                                                                                                                                                                                                                                         -----ATCCGCC
                                                                                                                    roAlaProGlyLeuAlaSerGlyAspArgProProSerValSerSerValHisSerGluG
                                              33331 GCTCAGCACCTCCTTCCAGGAGGAGGAGCAGTTCCTCGCCGTACCTCCCCCGGCTACGG
                                                                      ------Gly-LysAlaLysValSerGlyArgProSerSerArgLysAlaLysSerP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Novel recombinant DNA compounds comprising coding sequences for desceamine transferase gene of Streptomyces venezuelae, useful for
                                                                                                                                                                                                                                                                                                                                                                                                                                                          Cosmid pKOS023-27 containing S. venezuelae PKS gene cluster.
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98US-00073538.
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99US-01191339P.
99US-0134990P.
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06-MAY-1998;
28-MAY-1998;
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22-SEP-1998;
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                                                                    The present invention relates to recombinant DNA compounds that encode Streptomyces venezuelae (Sv) narbonolide polyketide synthases (PKSs). The recombinant PKSs are derived from narbonolide PKS and other genes involved in narbomycin and picromycin biosynthesis in recombinant host that results in reproduction of picromycin. Also disclosed are enzymes that results in the production of picromycin. Also disclosed are enzymes uch as those responsible for glycosylation and hydroxylation, (e.g. Cl2 hydroxylate (PicK), descendine biosynthesis, and desosaminyl transferase enzymes. The recombinant narbonolide, narbonolide derivatives, and enzymes. The recombinant satisfiction and as intermediates in the synthesis of compounds for pharmaceutical applications. The present sequence represents cosmid pKOSO23-27 containing S. venezuelae PKS gene
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ------150
substrate
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15163 TACCTCAAGCGCGTCACCCGAGCTGCAGCAGAACACCAGGCGTCTGCGCGAGATCGAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 GluSerHisSerTyrLeuProGluLeuGlyLysSerGluMetGluPheIleGluSerLys
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 desosamine transferase which transfers desosamine to
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ò		ପୁ	 26347 GCTGC-
QQ	25342 CCCGCAGGACGCGCGCTGGGACGTGGAGGGGCTGTACGACCCCGACCGGACGCGTCCGG 25401	Š	486 erTyrA
ò	181 ArgGlulleThrMetValGluGlnGlnIleSerLysLeuLysLysLysGlnGlnGln 199	අු	 26352C
q	25402 CAGGACGIACTGCCGGTCCGGCGGATTCCTGCACGACGCCGGCGACTTCGACGCCGACTT 25461	ð	506 lnGlnG
ò	200 LeuGluGluAlahlaLysProPro	qq	:: 26407 GGTGGA
Q O	CTTCGGGATCTCGCGCGAGGCCCTCGCCATGGAACCGCAGGAGCGAGC	à	526 lu
ò.	GlubrodiuLyskrovalserkrokio	qq	26466 GCGCGG
g	CACCGCGTGGGAGGCGATCGAGAGCGCGGGGCATCGACCGGCCCCTGAAAGGGGCGGGGGGGG	à	529 laGluL
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ò	270 IleLys1leAsnGlnAlaMetArgLysLysLeulleLeuTyrPheLysArgArgAsnHis 289	:	26706 GTACGC
q	25762 GICCTCGCTGGTCGCCTGCCTCGCCGTGCAGGCCCTCCGCAAGGCCGAGTGCGACAT 25821	ò	575 lnGly
ò	TyrAspGlnLeuMetGluAlaLeu	qq	26766 CCCTGC
g G		ò	595 laIle1
ò	310 GluLysLysValGluArgIleGluAsnAsnArpArgArgArgAlaLysGluSerLysVal 329	qa	26826 TCGTCT
qq		ò	613 er
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ò		q	26934 GCTAC
qq	ĠĠAGTCCTGCTGGTGGAGCGCCTGTCGGACGCCCGCCG	ò	650 ysAsn
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ò		qa	27016
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ŏ	ysGl	qq	27086 GTCGG
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ò	446 InHisproLygAsnPheGlyLeulleAlaSerPheLeuGluArgLysThrValAlaGLUC 466 ::: :::	qu	27145 CGACG
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466	ysValLeuTyrTyrTyrLeuThrLysLysAsnGluAsnTyrLysSerLeuValArgArgS 486
7	 GCTGC
486	erTyrargargalyLysserGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnG
506	InglnGlnGlnGlnProMetProArgSerSerGlnGluGluLysAspGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLysGluLy
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WPI; 2000-072618/06.
P-PSDB; AAK67201, AAY67202, AAY67203, AAY67204, AAY67205, AAY67207,
AAY67208, AAY67211.
                                                                                                                             Narbonolide polyketide synthase; PKS; cosmid pKOS023-27; ketolide; antibiotic production; narbomycin; picromycin; ds.
                                                                                                            Recombinant cosmid pKOS023-27 containing S. venezuelae PKS genes.
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/product= "PicB"
/note= "Contains typeII thioesterase domain"
34863. .36011
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                                                                                                                                                                                                                                                                                                                                                                                         product= "PICCII"
note= "4-keto-6-deoxyglucose isomerase"
                                                                                                                                                                                                                'note= "Narbonolide synthase subunit 1"
                                                                                                                                                                                                                                                  'note= "Narbonolide synthase subunit
55133. .29821
                                                                                                                                                                                                                                                                                                                           note= "Narbonolide synthase subunit 13961. .34806
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                                                                                                                                                                                                                                                                                                                                                                                                                                                         /*tag= h
product= "PICCVI"
/note= "3-amino dimethyltransferase"
                                                                                                                                                                                                                                                                                                                                                                                                                                      note= "Desosaminyl transferase'
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Mcdaniel R,
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70. .13725
/*tag= a
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                                                                                                                                                                                                                                                                                                                                                                                                                              "PICCIII"
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product= "PICAII"
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product= "PICAIV"
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'product= "PICAI"
                                                     AAZS6001 standard; DNA; 38506 BP
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98US-00141908.
98US-0100880P.
99US-0119139P.
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           33655 CCTGTTCGCGGG 33666
ProLeuAlaGly 2493
                                                                                          (first entry)
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/product=
                                                                                                                                                         Streptomyces venezuelae.
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28-AUG-1998;
22-SEP-1998;
08-FEB-1999;
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                                                                                          23-MAR-2000
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This is the recombinant cosmid pKOS023-27 DNA sequence which contains a Streptomyces venezuelae DNA insert. The cosmid contains open reading frames which encode the various modules of the narbonolide polyketide captures (PKS). The invention relates to recombinant DNA containing a coding sequence for a narbonolide PKS. Polyketides are compounds and coding sequence for a narbonolide PKS. Polyketides are compounds and coding sequence for a narbonolide PKS. Polyketides are compounds and coding module, six extender modules, are responsible for the production of many antibiotics including picromycin. The narbonolide PKS consists of a loading module, six extender modules, and two thioester domains. Four proteins make up the narbonolide PKS (PICAI, PICAII, PICAII and PICAV). PICAI includes extender modules and extender modules and and extender module 5 and contains and produces and two finesterase domain. The second type II thioesterase domain. The second type II thioesterase domain is found on the PICB protein. The cucleotide sequences encoding all of these proteins can be isolated in recombinant form from the recombinant cosmid pKOS023-27. Narbonolide is desosaminylated in S. venezuelate to yield narbomycin, the desosaminylater bloss the invention is used to express, in transferase enzyme is required for this conversion, and the desosamine DNA of the invention is used to express, in transferade captures.
New recombinant DNA encoding a domain of narbonolide polyketide synthase, for production of ketolide antibiotics.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     polyketide antibiotics or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             narbonolide (or its derivatives) or other ketolides (particularly hybrids), which may then be converted (e.g. by other enzymes recombinantly expressed in the same hosts) to polyketide antibiotitheir intermediates. The antibiotics are useful in human or veterit
                                                                                     Example 2; Page 16-27; 98pp; English.
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Sequence 38506 BP; 4914 A; 15118 C; 13444 G; 5030 T; 0 U; 0 Other;

			72	2494	98	2500	106	2505	123	25103	138	25163	142	2522	150	25282
			LeuSerProGlySerIleIleGlnProGlnArgArgArgProSerLeuLeuSerGluPhe 72	CTCGCGCCGCCGCCGCGGCGGCCGCCGCCCCCGGCTCAGGGCGCGCGC	-LeuHisLeuArgPro	GGCCCTGGGGGACGACGACGACGACCTGGACGAGGCGTCCGACGACGACCTC	GluSerHisSerTyrLeuProGluLeuGlyLysSerGluMetGluPhelleGluSerLys	-ctictccttcatcgacaagctgggcgactccgactictgacctgcccg	ProLeu	ccgccaccaccaccaccccccccccacacaca	-LeuAlaThrGlyGlnProAlaGlySerGluAspLeuThrLysAsp	GACAAGCTCCGCGAC	ArgSerLeuThr	25163 TACCTCAAGCGCGTCACCGCCGAGCTGCAGCAGAACACCAGGCGTCTGCGCGAGATCGAG	OzdPro	GACGCACGCACGACCGGTCGCGATCGTGGGCATGGCCTGCCGCCCTGCCGGGCGGTGTC
•	38506 697 697 1150 1061		ArgProSer			SGACGAGGC	31uMetGlu	CGACTCCGA	ArgProSez	CCACCAGC	SlySerGlu	AACAACGAA		ACCAGGCGT	1	SCCTGCCGC
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	1.73e-09 00.00 31.71# 21.69# 4.54#	US-09-522-753-5 (1-2517) x AAZ56001 (1	oGlySerileileGln	GCCGCGGGGGGGT	GlnProGlyAsnGluArgSerGlnGlu	 	sSerTyrLeuProGlu	-cricricricarca	107 ArgProArgLeuGluLeuLeuProAspProLeuLeuArgProSerProLeu	555 555	LeuAlaThr	25103 ACACGGAACGGACAGGCGAGAACGGAGGCCATGGCGAACAACAACGAAGACAAGCTCCGCGAC		GCGCGTCACCGCCGAG	GlyLysLeuGluProValSer	gcacgagccggragcg
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163

-AspProGluLeuGluLeu---Val

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1888 rAlavalGl ::: 31799 GTCACCTTC	AlavaiglubroserLyskrointvalledargserintserintserint	a è	32698 CGA 2238 qSe
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1947 gProGlu- 31946 ACCCTCG	<pre>gprogluArgProArgAlaAspThrGlyHisAlaPheLeuAlaLysProProAl 1965 </pre>	g :	32865 TGA
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2163 euTyrSe 	2163 euTyrSerPheProGLyAlaberCyBrrovalLeumsbreumignigtoriosernebr 2259 22539 concentrate the control of the cont	ය .	
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oalaproglyLeualaserGlyAspArgProbroserValSerSerValHisserGluG 2437 regargegerearcegergeregeregegerecareagaegaegeges 33270 CTCAGCACCTCCTTCCAGGAGGGGGGACTTCCTCGCCGTACCTCTCCCCGGCTACGG 33390 ||||::: ||||::||||| arginecegecegeageceges 33150 pGluAspArgProSerSerAlaGlySerThrProPheProTyrAsnProLeuileMetA 2470 gLeuGlnalaGlyValMetAlaSerProProProProGlyLeu-ProAlaGlySerGly 2489 ||| :::|||||||| ::: |-----AGCGGAACTGGGCC-GTCGCCGAGCCGTCGGATCACGAGCGGAGGA 33036 y------AspCysAsnArgArg-----ThrProLeuThrAsnArgValT 2450 GAAGAGGTCCC------CGTCGACCGCCCGCTGCGGGAGATCGGCTTCGACTCGC 32864 --LeuGluAjaileIleArgLysAjaLeuMetGlyLysTyrAspGlnTrpGluGjuSe 2360 roProLeu-----SerAlaAsnAlaPheAsnProLeuAsnAlaSerAlaSer---- 2376 -----CCGACCTCCCGACGTACGCCTTCCAGCA------CCGCTCGTACTGGAT 32637 accecacidateceaacidadacaceceaacacearteceaacacacaadacearcac 32697 erLysSerproGlyAsnThrSerGlnProProAlaPhePheSerLysLeuThrGluSe 2278 snSerAlaMetValLysSerLysLysGlnGluIleAsnLysLysLys---LeuAsnThrHi 2297 :: accecetedactrececaacesetaacesetaacesetrecesetreceseta 32924 snarg---AsnGluProGluTyrAsnIleSerGlnProGlyThrGluIlePheAsnMe 2316 rohlalleThrGlyThrGlyLeuMetThrTyrArgSer-----GlnAlaValGl 2333 lluHisAlaSerThrAsnMetGly----- 2341 sSerGluGlyGlyLysArgSerProGluProAsnLysThrSerValLeuGlyGly-Gl 2218 uAspGly11eGluProValSerProProGluGlyMetThrGluProGlyHisSerAr 2238 eralayalTyrProLeuLeuTyrargAspGlyGluGlnThrGluProSerargMetGl 2258 -----Gly-LysAlaLysValSerGlyArgProSerSerArgLysAlaLysSerP Ser-----AspHisThrLeuThrSerProGlyGlyGly------BspHisThrLeuThrSerProGlyGlyGly------

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27559 CCATCGCGTGGGCTTCGCCCCCCGCGTGGAACCTTGGCGGTTGACGGCTTCACCCACTT 27618	814SerProSerAlaProProValValProLysGluGluLysGluGluGluThrAlaA 833 27619 27619 CATCGAGGTCAGCGCCCACCTCACCATGACCCTCCCGAGACCGTCACGGCCT 27678	833 laAlaProProValGluGluGluGluGluGluGln		27739 CIGGGCCAACGGCCICACCAICGACIGGGCCCAICCICCCAACCGCAACCGGCCACCA 27798	867 hrgluglualaglugluglyProAlaLy8	GlyLysAspAlaGluAla	27859 CACCAGCGCCGACGACGACTGGCGTTACCGCGTCGAGTGGAAGCCGCTGACGGCCTCCGG 27918	 rcgrcgccgr	006 006	27955 CGGGAGCGAGCCAGAAGCCGAGCTGCTGGGCGCGCTGAAGGCCGCGGGAGCGGAGGTCGA 28014		28015 CGTACTGGAAGCCGGGGGGACGACGACGTGAGGCCCTCGCCCGGCTCACCGCACT 28074	914 921 		936		euLeuThrProThrGlyAspP 951		951 roArgAlaAsnAlaSerProGlnLysProLeuAspLeuLysGlnLeuLysGlnArgAlaA 971			28309 COACCICCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCC	28357 CGCACTCTCCGGCGCCACGGCGAGGACCAGATCGCC		28417 CCGCCGCCTCGCCGCGCACCCCTCCACGGACGTCGGCCCACCCCACCGCGACTGGCAG	1013	28477 CGGCACCGTCCTCATCACCGGCGGCACCGGAGCCCTCGGCACGCCACGCTGGAT	1030AlaAspLysGluAlaPheAlaAlaGluAlaGlnLysLeuProGlyA 1045	28537 GGCCCACCACGGAGCCGAACACCTCCTCGTCAGCCGCAGCGGCGAACAAGCCCCCGG	. 1045 spProProCysTrpThrSerGlyLeuProPheProValProProArgGluValIleLysA 1065
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97	AACTCACCGCCGAACTCACCGCATCGGGCGCCCGCGTCAC 28644
1065 laSerProHisAlaProAspProSe : 28645 CATCGCCGCCTGCGACGTCGCCGAC	87
1085 roLeuGlyLeuHisAspThrAlaAr 28705 CG	aArgProValLeubroArgProProThrIleSerAsnP 1105
1105 roproproLeuIleSerSerAla 28735 CACCGCCGCACCGGCGCGAT	roProproLeuIleSerSerAla
. 1116 erValLeuGluArgGlnIleGlyAl :: :: 28795 CATCCTGGGGGGAAGACGAGGGGG	ervalLeuGluArgGlnIleGlyAla1leSerGlnGlyMetServalGlnLeuHisValP 1136 :: :: carccrggggggggaAgacgaggggggggggggacgccgggacrcc 28854
1136 roTyrSerGluHisAlaLysAlaPr :::::: 28855 GCTGGACGCTTCGTCCTCTACTCC	roTyrSerGluHisAlaLysAlaProValGlyPro
1148 alThrMetGlyLeuProLeuPro ::: 28915 CGTCTACGCGGCGACGACGCCAA	alThrMetGlyLeuProLeuProMetAspProLysLysLeuAlaProPheS 1165 :::
1165 erGlyvalLysGlnGluGlnLeuSe ::::: 28971GGGGGAGACGGCGACTCGGT	erGlyvalbysGlnGlnGlnLeuSerProArgGlyGlnAlaGlyProProGluSerL 1184
1184 euGlyvalProThrAlaGludluAl	euglyvalprothralaglnglualaserValLeu
1196	A 1196
29086 CGCCCTGGACGAACTGGCCAAGGC	CGCCCTGGACGAACTGGCCAAGGCCCTGAGCCACGACGAGACCTTCGTCGCCGTGGCCGA 29145
1196 rgGlyThrAlaLeuGlySerValP: 29146 rGrCGACTGGGAGCGGTTCGCG-C	rgGlyThralaLeuGlySerValProGlyGlyBerIleThrIvs1210
1211GlylleProSerThrArg :: 29205 ACGGCGTCCCGGAGGCCCGGCAGG	Glylleproserthrarg
1220	spSerAlalleThrTyrArgGly
1231	HisGlyThrProAlaAspValLeuTyrLysGlyThrIleThrArgIleIleG 1248
1248	19G1uAspSerProSerArgLeuAspArgG1yArgG1uAspSerLeuProLysG1yHisV 1268
1268	allleTyrGluGlyLysLysGlyHisValLeuSerTyrGluGlyGlyMetSerValThrG 1288
1288	IncysserLysGluAspGlyArgSerSerGlyProp 1301
2 1	tGluGlyArg 131 GGAGGGGGG 296
1318 ValGlyArgAlalleSerSerAle	valGlyArgAlaileSerSerAlaSerIleGluGlyLeuMetGlyArgAlaileProPro 1337

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            New recombinant pick hydroxylase gene of Streptomyces venezuelae useful for converting ketolides to antibiotics and as antibiotics and intermediates in the synthesis of compounds with pharmaceutical value.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    synthase; polyketide synthase gene; narbonolide polyketide;
C12-hydroxyalse; picK; desosamine biosynthesis;
transferase enzyme; ketolide; beta-glucosidase enzyme;
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                        ---AspCysAsnArgArg----
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06-MAY-1998;
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20-MAY-1999
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                                                                                                                                                                                                                                                                                                                                                                                           GluSerHisSerTyrLeuProGluLeuGlyLysSerGluMetGluPheIleGluSerLys 106
These compounds are also useful for increasing the antibiotic activity of a compound relative to the unhydroxylated compound. The recombinant host cells are useful as genetic systems that allow rapid engineering of the narbonolide polyketide synthase. These would be valuable for creating novel ketolide analogs for pharmaceutical applications
                                                                                                                                                                                                                                                                                                                                    86
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24947 GGCCTGGGGGACGACGGCGACCACCGACCTGGACGAGGTCTCGACGACGACCTCTA
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                                                                                                                                                                                                                                                                                                                                                                                                                                                       107 ArgProArgLeuGluLeuLeuProAspProLeuLeuArgProSerProLeu----
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Matches:
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                                                                                                                                                                                                                                                                                                                                         73 GlnProGlyAsnGluArgSerGlnGlu--
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1815 CGGGGAAAACCGCACGACGTGAT.	Db 35270GGTCCCCCAGACACCCCCAGGTCCCCTTC 35297 Qy 2064 euThrHisAspLysGlyLeuProLysHisLeuGluGluLeuAspLysSerHisLeuGluG 2084

1156 tAspProLysLys 32666 cGACCGGCTGCGG	LABPROLYSLYS	op ox	1484 e
		λ d	1504
CGAGCCG	GGTTCCGGCGGTTCGGACGCGCGCCGCCGACCCTGGTGC 32773	'n	1524
12774 GGAGCCGGAGGCGTCGATCGA-CGACCTGGACGCCGAGGCCTGATCCGGATGGCTCTCG		g è	33651
1208IleThrLysGlylleProSerThrArgValProSerAspSe	LysGlylleProSerThrArgValProSerAspSe 1221 	G &	33699
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	ACCCCATCC32929	<u> </u>	33/49
		qa qa	33809
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1261 pSerLeuFroLysGlyHisVallielYrGiuGlyLySLySGlyHisValLeuSerlYrGi 32889 GCGCTCTCAAGGAGA-ACGAAGAACTCCGAAAAAAGAGCCGTCGCCGGGCCGAC	ySLYSGIYHISVAILGUSEIIYIGI 1201 	ΩD	33869
1281 uGlyGlyMetSerValThrGlnCysSerLysGluAspGlyArgSerSerGl 	ysGluAspGlyArgSerSerSerGl 1299	රු අ	33929
33045 cgrcddcagaadccargacgarcgrcdgcargacrgccggrrcgcGGGCGGAarcCGG		λ	1631
1299 yProproHisGluThralaalaProLysArgThrTyrAspMetMetGluGl 	ArgihrTyrAspMetMetGluGl 1316 :: CAAGGACCTGGTCTCCGAGGTACCG 33164	qq	33978
		ò	1651
		8 8	34026
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33225 ACGTACGTCCGCAACGCCGCGTTCCT		ò	1691
1354 erileThrGlnGlylle-ProArgSerTyrValGluAlaGluAspTyrLeuArgArg	luAlaGluAspTyrLeuArgArg 1373	a	34094
		ò	1711
1374 GluAlalysLeuLeuLysArgGluGlyThrProProProProProProSecArgAspleu	OProProProSerArgAspleu 1393	qa	34154
GAGGCCCTCGCCATG	0000	ò	1726
ThrGluAla		ପୁ	34214
		λο	1729
AlaHisGluGiyLeuValAlaThrValLysGl		qa	34274
		ΛO	1736
1430 ArgGluGluLeuArgHisThrProGluLeuProLeuAlaProArgProLeuLysGLuGLY	1449	qa	34334
5TACGCGCCGGACATCCGGGT		λõ	1753
	1.4	qa	34383
TACGICGICACCGGCAACTCCTCO		ò	1773
1469 sLysHisAspValArgSerL	-ValArgserLeuileGlySerProGlyArginrPn 1484	qa	34418
33537 CTGGAGGACCCGCCTGACCGTGGACACGCGT-	71971797177779	ò	1793

1484	eProProvalHisProLeuAspValMetAlaAspAlaArgAlaLeuGluArgAlaCysTy 1504
1504	rGluGluSerLeuLysSerArgProGlyThrAlaSerSerSerGlyGlySerIleAla-A 1524
1524	rgGlyAlaProValileValProGluLeuGlyLysProArgGlnSerProLeuThrTyrG 1544
1544	luaspHisGlyAlaProPheAlaGlyHisLeuProArgGlySerProValThrMetArgG 1564
1564	IUProThrProArgLeuGlnGluGlySerLeuSerSerSerLysAlaSerGlnAspArg
1584	
1602	GluHisHisProHisProlleSerProTyrGluHisLeuLeuArgGlyVal 1618
1619	
1631 33978	
1651	uProArgHisLeuAlaProAsnProThrTyrProHisLeuTyrProProTyrLeuIleAr 1671
1671	gGlyTyrProAspThrAlaAlaLeuGluAsnArgGlnThrIleIleAsnAspTyrIleTh 1691 GGCAGCCG
1691	rSerGlnGlnMetHisHisAssnThr. GCACACGCAGGCCGCTTCGGGTGTC
1711 34154	
72	
1729	ArgGlyIleIleAspLeuSerGl 1736
34274	GAGGA 343
1736	6 nValProHisLeuProValLeuValProProThrProGlyThrProAlaTh 1753 :::
1753	3 ralaMetAspArgLeuAlaTyrLeuProThrAlaProGlnProPheSerSerArgHisSe 1773
1773	3 rSerSerProLeuSerProGlyGlyProThrHisLeuThrLysProThrThrThrThrSerSe 1793
1793	3 rSerGluArgGluArgAspArgAspArgGluArgAspArgAspArgGluArgGluLysSe 1813

Db 31627 CCACCCAACTCACCGCGAACTCACGGCGCGCGCGCGCGCTCACCATCGCGCCTGCG Qy 859 GluGluProValLysSerGluCysThrGluGluAlaGluGlyProAlaLysGlyLys	Qy 898 Db 31807	Cy 918 Db 31858	. da .	Oy 954 Db 31939	Oy Db 31	Db 32041 GCGACGGCGGGGCGCCGACGACGCGTACTGGCAGGCTCGCGCGCATCCGTCCG	Cy 990AlaProThrLysProAlaProProAlaProProProProGlnAsnLeuGlnDroGlu	Oy 1009 SerAspAlaProGlnGlnProGlySerSerProArgGlyLysSerArgSerProAlaPro	Oy 1029 ProAlaAspLysGluAlaPheAlaAlaGluAlaGluLysLeuProGlyAspPro	Qy	Db 32323 CGGCGCTGGCGATCACCGCGCTCCCCCGAGCCCGAGCGCCGGCCGGCGCTCCTRA	Qy 1081 GlyHisProLeuGlyLeuHisAspThrAlaArgProValLeuProArgProProPro	Oy 1101 ThrileSerAsnProProProLeulleSerSerAlaLys-HisProSerVal 12432GTGGCCCCGGCCGTCACCGAGCTCGGCTTCGACTCGCTTCACCGGCTTCGACTCGCTGACGGCCGTTCACCGACTCGACTCGCTTCACCGGCTTCACCGGCTTCACCGGCCGTTCACCGGCCGTTCACCGGCCGTTCACCGGCCGTTCACCGGCCGTTCACCGGCCGG	Qy 1118LeuGluArgGlnIleGlyAlaIle	Oy 1126SerGlnGlyMetSerValGlnLeuHisValProTyrSerGluHisAl	31626 Qy 1141 aLysAlaProValGlyProValThrMetGlyLeuProLeuProMe
	30879 TCCGCGCCCACCACCGACGACTGGCGTTACCGCGTCGAGTGGAAGCCG 30932 570 Ly8ThralaAsnSerGlnGlyArgArgLy8GlyArgIleThrargSerMetalaAsnGlu 589 101	590 AlaAsnSerGluGluAlaIleThrProGlnGlnSerAlaGluLeuAlaSerMetGluLeu 609 30993 CCAGAAGCCGAGCTGCTGGGCGCGTGAAGGCCGCGGGAGGCGGAGGTCGACGTA 31046	610 AsnGluSerSerArgTrpThrGluGluGluMetGluThrAlaLysLysGlyLeuLeuGlu 629 31047 CTGGAAGCCGGGGGGGGACGACGACGACGTGAGGCCTCGCCGCCCGGCTCACCGCACTGACG 31106	630 HisGlyArgAsnTrpSerAlaIleAlaArgMetValGlySerLysThrValSerGlnCys 649	650 LysAsnPheTyrPheAsnTyrLysLysArgGlnAsnLeuAspGlulleLeuGlnGlnHis 669 31140	670 LysLeuLysMetGluLysGluArgAsnAlaArgArgLysLysLysLysLysAlaPro 687 :::		GluAlaGluAlaLeuHisAla	SerGlyAsnGluValProArgGlyGluCysSerGlyProAlaThrValAsnAsnSerSer	747 AspThrGluSerIleProSerProHisThrGluAlaAlaLysAspThrGlyGlnAsnGly 766 11:: :::		ProThrProProArgArgThrSerArgAlaProIleGluProThrPro	31387 CHCITCCGGCGCCCCACCGGCGCGCGCCCCCCCCCCCCCCCC			

The invention relates to an isolated and purified nucleic acid segment comprising a desosamine biosynthetic gene cluster, a fragment or its biologically active variant, where the nucleic acid sequence is not derived from the eryc gene cluster of Saccharopolyspora erythreae or Streptomyces antibioticus. The invention also relates to a macrolide biosynthetic gene cluster or fragments thereof. The macrolide compounds. Recombinant or augmented cells compound the desosamine compounds. Recombinant or augmented cells compound the desosamine and/or macrolide biosynthetic gene clusters are useful for the production of biologically active macrolides. The macrolide biosynthetic gene clusters are useful for the production of biologically active macrolides. The macrolide biosynthetic gene clusters are useful for the production of biologically active macrolides. The macrolide biosynthetic gene clusters are useful for synthesis of methywycii, pkromycin, neomethymycin, and clusters are useful for synthesis of methywycii, pkromycin, neomethymycin, and active active for an arbomycin. The alternative termination of polyketide synthesis may be useful to prepare novel antibiotics and polyhydroxyalkanoate (PHA) monomer synthases or to prepare biologically active agents, case biopolymers, e.g., in packaging or biomedical applications, to engineer PHA monomer synthases or to prepare biologically active agents, chronic obstructive pulmonary disease as well as other diseases involving respiratory inflammation, cholesterol-lowering agents or macrolide-based chronic obstructive pulmonary disease as well as other diseases involving respiratory pathogens, as well as viral parasitic pathogens, or as crop respiratory pathogens, as well as viral parasitic pathogens, or as crop respiratory pathogens, as well as viral parasitic pathogens, or as crop respiratory pathogens, as well as viral parasitic pathogens, or as crop protection agents (e.g., funglides or insecticides) via expression of polyketides in plants. The present sequence represents th ¥8888888888888888888888888888888888

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> Other; 0 ņ Sequence 37948 BP; 4901 A; 14718 C; 13311 G; 5018 T; 0

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> Length: Matches: Conservative: Mismatches: Indels: Gaps: 1.71e-09 600.00 33.91% 22.97% 4.54% Similarity: Percent Similarity Best Local Similari Alignment Scores: Best Local S Query Match:

eccesesesesertrestrircceseccases services escaped esca ------ccgaactccrrdacactcaccddacrrccrrccarrgcccaa 29942 29992 29797 29673 29694 147 127 uSerHisSerTyrLeuProGluLeuGlyLysSerGluMetGluPheIleGluSerLysAr 107 sleuserProglySerIlelleGlnProGlnArgArg-------ArgProse 67 87 22 oValSerProProSerProProHisThrAspProGluLeuGluLeuValProProArgLe yGlnProAlaGlySerGluAspieuThrLysAspArgSerLeuThrGlyLysLeuGluPr 29943 TGCGAGACGCGCTCTCCCGC-TACGTCGACTGGTCTCTTGAAGCCGTC-----GT gProArgLeuGluLeuLeuProAspProLeuLeuArgProSerProLeuLeuAlaThrGl ArgAlaThrGluproArgTyrProProHisSerLeuSerTyrProValGlnIleAlaArg 5D-----rleuleuserGluPheGlnProGlyAsnGluArgSerGlnGluLeuHisLeuArgProGl ThrHisThrAspVal-GlyLeuLeuGluTyrGlnHisHisSerArgAspTyrAlaSerHi 29755 CGGTCGCGGTCGCGACAGCCGGAGGCACTGCGGGA------29626 connencescococococococococon con contra con contra c (1-37948)US-09-522-753-5 (1-2517) x AAZ87285 29674 ACGCGGACGGTCGTACGGA-107 127 25 67 147 33 29842 29798 g 엄 유 ઠ 유 ð 요 8 à 원 ò ò 셤 ò ò

--GCCCATCCTCCCCACCGCAACC 30818 30640 cccactrcarcaagarcagcdccaccccarcarcarcarcarcarcarcaagaccgrca 30699 ----cgaadccregeccaaceccrcaccarceac 30791 ::: ::: |||| gcttcgcccccccccagagaccttggcggttgacgcc-------TTCA 30639 ::: :::::||| ----Gacccagarcaagaacrcgcaccacaccacaagccaacgccacaaa30408 -- Geodalatre de cece ce ca de contra de contr 30098 cacccccaeaccercesceactecaeceaesesaaarcecececeaeceaececececes 30211 TCGCCGCCAC----CTCGCCGGCAAGGCGGGATGATCTCCCTCGCCCTCGACGAGG 30264 410 leAsnMetAsnGlyLeuMetAlaAspProMetLysValTyrLysAspArgGlnValMetA 430 snMetTrpSerGluGlnGluLysGluThrPheArgGluLysPheMetGlnHisProLysA 450 510 lnProMetProArgSerSerGlnGluGluLysAspGluLysGluLysGluLysGluLysGluAlaG 530 239 slysleulleleuTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPh 298 206 uSerLysGluGluLeuIleGlnAsnMetAspArgValAspArgGluIleThrMetValGl 187 snPheGlyLeulleAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrT yrTyrLeuThrLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArgA 370 isGluValSerGluIleIleAspGlYLeuSerGluGlnGluAsnLeuGluLysGlnMetA ccgccrccacccrccaccacaaacaaga-agaccagaagcgrcragrcac----snAsnProArgArgAlaLysGluSerLysValArgGluTyrTyrGluLysGlnPheP roglulleArgLysGlnArgGluLeuGlnGluArgMetGlnSerArgValGlyGlnArgG TCATCCCGGTCGACTACGCCTCCCACAGCCGGCAGGTCGAGATCATCGAGAAGGAGCTGG -----MetSerAlaAlaArgSerGluH rgGlnLeuAlaValIleProProMetLeuTyrAspAlaAspGlnGlnArgIleLysPheI leGluSerLysHisArgSerLeuValGlnIleIleTyrAspGluAsnArgLysLysAlaG 239 lualaalaHisargIleLeuGluGlyLeuGlyProGlnValGluLeu-ProLeuTyran eCysGln-ArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGluArgIleGluA GlnProSer-AspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLy uGlnGlnIleSer-LysteuLys---LystysGlnGlnGlnLeuGluGluGluAlaAlaL ccgacaggaacccgcccacccacgcrcaaccccacgac 358 lySerGlyLeuSer------ysProProGluProGluLysProVal--30469 CCGAGGTCCTC----------CTCACTCGC-470 390 430 30700 450 30752 30792 219 259 278 318 338 30480 30751 30265 298 30409 30598 30353 167 187 206 29993 g යු දු g ò 임 ò . g ò 셤 ð 임 ò ò

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	31647 ATCGAGATGGCTCTCGGCCCCGTAACACTGACCGGACCG	Qy 2482 roGlyLeuProAlaGlySerGlyProLeuAlaGlyProHis 2495 +	S. venezuelae pik (n Desosamine biosynthe neomethymycin; narbo biopolymer; antibiot chronic obstructive hypercholesterolaemi Streptomyces venezue WO200000620-A2.	06-JAN-200 25-JUN-199 26-JUN-199 (MINU) UD Sherman DF WPI; 2000- P-PSDB; 'AA	synthesis of methymycin and pikromycin. Claim 13; Page 299-315; 438pp; English.
UGluLeuAspLysSerHisLeuGluGlyGluLeuArgPro	GGGGTCTGGGGCAGCCAGGCGTCTACGGGGCCCAACGC	182 spleuTyrLeuProProAsp		2267 InproproAlaPhePheSerLysLeuThrGluSerAsnSerAlaMetValLysSerL 2286	31405 CGCACCTCCACGAGGCGTACCTCGCCCGAGCCGGCCCCGACGGACTGGG 31458 2343 lualaileileargLysAlaLeuMetGlyLysTyraspGlnTrpGluGluSerProProL 2363

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ò		쉱	29545
අු	GGTGGCGTTCGTCTTCCCC	δ	1832 uGln
δ		q	::: 29592 GAAG
qq	28695 GGCCAGGGCACGCAGGGCCCGGCATGGGCGCCGAACTCCTTGACAGCTCACCGGAGTTC 28754	ò	1852 aSer
δ	1534	qa	::: 29652 CACC
qa	28755 GCTGCCTCGATGCCGAATGCGAGACCGCGTCTCCCGCTACGTCGACTGGTCTT 28811	ò	1872 nGln
δλ	1547 yAlaProPheAlaGlyHisLeuProArgGlySerProValThrMetAr 1563	q	29679 TTCC
qq	28812 GAAGCCGTCCGACAGGAACCCGGCGCACCCACGCTCGACGTCGACGTCGTC 28868	ò	1892 oSer
ò		q	::: 29721 AGCG
qu		ò	1912 rPhe
ò		q	29773
q		ò	1927
ò	1593 aLysSerProHisSerThrValProGluHisHisProHisProIleSerProTyrGLuHi 1613	qu	29829 GCCG
q		ò	1928
δλ		Q	29889 GCGG
qq	ATCTCCTCGCCTCGA	ò	1943 oArg
ò		අු	29940 GGCG
qq	29090 CGAGGCGGCCGTCCTGAAGCGACTGAGCGACTTCGACGGACTCTCCGTCGCCGCCGTCAA 29149	* &	1956 yHis
ò	1649 rTyrLeuProArgHisLeuAlaProAsnProThrTyrProHi 1663	q	30000 GTGC
qg	29150 -cGGCCCCACCGCCACCGCCTCCGCCGACCCCAGATCGAGAACTCGCCCGCA 29208	ò	1967
ζö		q	30060 GTCT
q		δ	1970 uPro
ò	yrileThrSerGlnGlnMetH	q	30120 GGCC
qq	29251 ACTACGCCTCCCACAGCCGGCAGGTCGAGATCAGG	ò	1981 oArg
ò	1696 isHisAsnThrAlaThrAlaMetAlaGlnArgAlaAspMetLeuArgGlyLeuSerProA 1716	q) 30180 CAGC
qq	29289	ò	1997 hrPr
ò	1716 rgGluSerSerLeuAlaLeuAsnTyrAlaAlaGlyProArgGlyIleIleAspLeuSerG 1736 	qq	30240 GACC
đ	29325	à	2013 laPr
ò	hrA	q	30294 CCC
đ	29326 AGGCTCCGCACGTGCCGTTCTTCTCCACCCTCGAAGGCACCTGGATCACCGAGC 29379	ò	2022
ò	1754 laMetAspArgLeuAlaTyrLeuProThr-AlaProGlnProPheSerSerArgHisSer 1773	Ω	30354 GGCC
ф		ò	2040 erLe
ò	.euSerProGlyGlyProThrHisLeuThrLysProThrThrSer	. ପ୍ର	30387 CGC
q		λō	2057 erP
ò		g	30447 GCC
qq	29485 TCAGGGCCCACCCGTCCTCACCATGACCTCCCGAGACCGTCACCGGCCTCGGCACCC 29544		

1812	sSerIleLeuThrSerThrThrValGluHisAlaProlleTrpArgFroGlyThrGl 18	832 9591
1832	uGlnSerSerGlySerSerGlySerSerGlyGlyGlyGlyGlyGlyGlyGlyGerSerBerArgProAl 1 :::::	852 9651
1852	aserHisSerHisAlaHisGlnHisSerProlleSerProArgThrGlnAspAlaLeuGl ::: CACCACCCCGAGCTCCCCACCTACGCC	
1872	nGlnArgProSerValLeuHisAsnThrGlyMetLysGlyIleIleThrAlaValGluPr 18	892 9720
1892	oSerLysProThrValLeuArgSerThrSerThrSerBroValArgProAlaAlaTh 19 ::: AGCGCCGCCGACTGGCGTTACCGCGTCGAGTGGAAGCCGCTGACGGCCT 29	912 9772
1912	rPheproProAlaThrHisCysProLeuGlyGlyThrLeuAspGly	927 9828
1927		927
29829	GCCGAGCTGCTGGGCGCGCTGAAGGCCGCGGGAGCTCGACGTACTGGAAGCCGGG 2	9888
1928	ValtyrProThrLeuMetGluBroValLeuLeuProLysGluAlaPr 1	943
1943	OARGVALALAARGProGluArgProArgAlaAspThr	956
1956	YHisAlaPheLeuAlaLysProProAlaArg	966
1967	SerGlyLeuGl 1	. 026.
30060	ACCCCCGCCGACCCGGCCATGCTCTGGGCCTC 3	0110
30120	0 uProAlaSerSerPro	1981
1981	3 1	1997
30240	rcccccccc 3	2013
2013	3 laProProAlaSerAlaSerAspPro	2021 30353
30354	2HisArgGluLysThrGlnSerLysProPheSerIleGlnGluLeuGluLeuHugs 2	2040 30386
30387	erLeuGlyTyrHisGlySerSerTyrSerProGluGlyValGluProVals 2 ::	30446
30447	7 erProValSerSerProSerLeuThrHisAspLysGlyLeuProLysHisLeuG 2	2075 30506

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Disclosure; Fig 31; 438pp; English
                                                    Liu H,
                                     WO200000620-A2
                                           25-JUN-1999;
                                              26-JUN-1998;
                                                    Sherman DH,
                                        06-JAN-2000.
             Key
                                CDS
                  SDS
                       S
                           SGS
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Desosamine biosynthesis; macrolide; polyketide; methymycin; pikromycin; neomethymycin; narbomycin; polyhydroxyalkanoate monomer synthase; biopolymer; antibiotic; chemotherapeutic; immunosuppressant; asthma; chronic obstructive pulmonary disease; respiratory inflammation; hypercholesterolaemia; crop protection agent; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Desosamine and macrolide biosynthetic gene clusters, useful for, e \cdot g \cdot synthesis of methymycin and pikromycin.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1782. .35822

/*tag= d

/product= "Pik gene cluster protein #4 (AAY77203)"

/*tag= e

/*tag= e

/product= "Pik gene cluster protein #4 (AAY80997)"
                                                                                                                                                                                                                                                                                                                                                                                                                                                     'product= "Pik gene cluster protein #3 (AAY77202)"
                                                                                                                                                                                                                                                                                              "Pik gene cluster protein #1 (AAY77200)"
                                                                                                                                                                                                                                                                                                                                                                           "Pik gene cluster protein #2 (AAY77201)"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             WPI; 2000-160679/14.
P-PSDB; AAY77200, AAY77201, AAY77202, AAY77203, AAY80997.
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                                                                                                                                                                                                                   Location/Qualifiers
1742. .15583
                                                                                                                                                                          Streptomyces venezuelae; ATCC15439
                                                                                                                                                                                                                                                                                                                               15688. .26907
/*tag= b
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> comprising a desosamine biosynthetic gene cluster, a fragment or its biologically active variant, where the nucleic acid sequence is not derived from the eryc gene cluster of Saccharopolyspora erythraea or Streptomyces antibioticus. The invention also relates to a macrolide biosynthetic gene cluster. Or fragments thereof. The macrolide pixtomycin, neomethymycin, narbomycin or a combination of these compounds. Recombinant or augmented cells comprising the desosamine of biologically active macrolides. The macrolide biosynthetic proteins of biologically active macrolides. The macrolide biosynthetic proteins are useful for synthesis of methymycin, pixtomycin, neomethymycin and are useful for synthesis of methymycin, pixtomycin, neomethymycin and useful to prepare novel antibiotics and polyhedroxyalkanoate (PHA) useful to prepare novel antibiotics and polyhydroxyalkanoate (PHA) as biopolywers, e.g., in packaging or biomedical applications, to engineer PHA monomer synthases or to prepare biologically active agents, such as chemotherapeutics, immunosuppressants, agents to treat asthma, chronic obstructive pulmonary disease as well as other diseases involving respiratory inflammation, cholesterol-lowering agents or macrolide-based antibiotics which are active against a variety of organisms, e.g., bacteria, including multi-drug resistant pneumococci and other respiratory pathogens, as well as viral parasitic pathogens, or as crop protection agents (e.g., fungicides or insecticides) via expression of polyketides in plants. The present sequence represents the macrolide biosynthetic gene cluster (pik) from Streptomyces venezuelae ATCC 15439, as given in figure 31. (Updated on 15-SEP-2003 to standardise OS field) The invention relates to an isolated and purified nucleic acid segment

equence 36778 BP; 4758 A; 14303 C; 12864 G; 4851 T; 0 U; 2 Other;	<pre>nment Scores: 1.67e-09 Length: 36778 . No.: 600.00</pre>		716 GluMetValGluGluAlaGluAlaLeuHisAlaSerGlyAsnGluVal 731	751	AACACCCGCTCACC 25	751 ePros	CCTCTCCGACGGCACCCTCGATGACGGCGGGG 2345	767	25 GAIGIGGAACACGIACIGCGICCCCCAACGCCCCCCCCCC	779 OPrOGIYPrOProThrProProArgArgThrSerArgAlarzolleGilleGilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollingilleTollin	85 TCGACGCCCGGCTACGACCIGGCAGCGITCGICATGITCICCI CCCCCCCCCCCCCCCCCCCCCCCCCCCCCC	799 aSerGlublaThrGlyAlaProintFrokIorrUblatatoriosi	21.6	 02 TGGCGCCGCCGGACAGCC	laGl 849	CTCAGCGAC	1 869	26104 -CGGCTGGCCCGTTCCGGGGCGACGCCATGGACAGCGAGCTGACCCTGTC 26153	888	acqcqqccarqcqccqqqqqqqqqqqqqqqqqqqqqqqq	88		903	037777	923 aAspGluValAspGluAlaGluGJVGJY	34 GGGCGAGGCGGACACACGGACCTCGC	C	6394 CGCGCACCTGCGGGACCTCGTCCGTACGCACGTGGCGACCGTCCTGGGALACGGCACCCC 2043.		957 oglnLysProLeuAspLeuLysGlnLeuLys	957 oglnLysProLeukspLeuLysGlnLeuLysGlnArgAla-AlaAlaIleProProIleG 97
Scor.	1.5	22-753-	716	m	5705	751	5765	76	282	77	288	79	ָּהְלָּהְ	009	N	605		610	9	615	88	621	90	627	92	633	m	39	ß		26454
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29998 29958 29958 30018 1436 1436 1436 1455 30138 30138 1467 1507 1507 151 1521 1521 1521 1524 1521 1521 1521 1535 1531 1609 30755 1649 1649 1649
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	29600 ĠAĠĠĠĠĠĠĠĠĠĠĠĊĠĊĠĠĠĊĊCGĠCCGCCCCGGCCGGGCGGG
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25959 —GGANCCTGTTCGTGCAGTTCAGCCGGCAGCGGGGGGGGGGGGGGGCGGGGGGGGGGG

613	erArgipphrglugluclumetGluThralaLysLysGlyLeuLeuGluH 630
63	AlaIleAlaArgMetVa
26934	BL 670
26951	
670	ysLeuLysMetGluLysGluArgAsnAlaArgArgLysLysLysLysLysLahaAlaA 690
D	710
27016	:::::: CGTGACCTTCGCTGTCATGGTC 27037
710	alserGlyAsnGluGluGluMetValGluGluAlaGluAlaLeuHisAlaSerGlyAsnG 730 ::::
73	luValpro
27086	GICGGCCACTCGCAGGGCGAGATCGCCGCCGCTGCGCCGGTGCACTCA-CCCTCGA 27144
741	hrvalAsnAsnSerSerAspThrGluSerIleProSerProHisThrGluAlaAlaL 760
160	ysAspThrG 763
27205	:: GGGCGGCATGATCTCCCTCGCCCTCGACGAGGCGGCCGTCCTGAAGCGACTGAGCGACTTT 27264
76	1yGlnAenGlyProLysProProAlaThrLeuGlyAlaAepGlyProProProGlyProP 783
27265	CGACGGACTCTCCGTCGCCGCCGTCAACGGCCCCACGGCCACCGTCGTCTCTCGG
7	rothrProProArgArgThrSerArgAlaProlleGluProThr 797
615/2	797
97279	GATCATCCCGGTCGACTACGCCTCCCACAGCCGGCAGGTCGAGA
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27439	
800	
27499	::: ::: caaaggcaccradarcaccaagccagracrcaacgacaccracragraccgcaaccracg
807	ThrProProProAlaProPro
27559	
814	SerProSerAlaProProValValProLysGluGluLysGluGluGluThrAlaA 833
27619	cardaagardaddacccccccccaddarcccccccagagaccgrcaccggccr 27
833	lablaProProvalGluGluGluGlyGlu
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27739	TIGHT TO THE TRANSPORT OF THE TRANSPORT OF THE TRANSPORT OF TRANSPORT

US-09-522-753-5 (1-2517) x ABS56090 (1-38506)

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-CCTTGGCACACGAT 23095
 ---TGTATCTCCTC 23044
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               narbonolide polyketide synthase (FKS) domain, and methods of producing polyketides by recombinant DNA technology. The recombinant DNA sequences are derived from Streptomyces venezuelae desosamine blosynthetic, desosaminyl transferase, beta-glucosidase, or pick (Cl2) hydroxylase genes. The method is useful for transforming a cell with a recombinant therefore for increasing the yield of a desosaminylated polyketide in a cell. The recombinant methods and materials are useful for expressing polyketides with significant antibiotic activity, derived in whole or in narbonolide PKS gene, and other genes involved in harbomycin and picromycin blosynthesis in recombinant host cells. The present sequence represents S. venezuelae DNA inserted into cosmid pKOS023-27 in the methods of the present invention
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    invention relates to recombinant DNA sequences encoding for
                                    2479 oProProGlyLeuProAlaGly---SerGlyProLeuAlaGlyProHisHisAla-T
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Preparation of polyketides by recombinant DNA technology, useful as antibiotics and as intermediates in the synthesis of pharmaceutical
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              C; 13446 G; 5030 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                               desosaminyl transferase gene; beta-glucosidase gene; antibiotic; pick hydroxylase gene; C12 hydroxylase gene; narbonolide; desosaminylated polyketide; narbomycin biosynthesis; mutant; picromycin biosynthesis; ds.
                                                                                                                                                                                                                                                                                                                                                                  Narbonolide polyketide synthase; PKS; desosamine biosynthetic
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Tang
                                                                                                                                              23096 Geccacadaracrercécecerérecerecaciár 23133
                                                                                                          rpAspGluGluProLysProLeuLeuCysSerGlnTyr 2510
                                                                                                                                                                                                                                                                                                                            S. venezuelae DNA inserted into cosmid pKOS023-27.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Mcdaniel R,
23000 TCAGCCTACCCCCAGCCCCCTCCTGCTGTCTCGC---
                                                                      23045 TGCACAGTTGGGTGCTCATGCCGGTGTTTCAGGCCCT
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Betlach M,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 38506 BP; 4914 A; 15116
                                                                                                                                                                                                                    ABS56090 standard; DNA; 38506 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (KOSA-) KOSAN BIOSCIENCES INC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            22-FEB-2001; 2001US-00793708.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        22-FEB-2002; 2002WO-US005642
                                                                                                                                                                                                                                                                                          (first entry)
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Synthetic.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Betlach MC,
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38506 700 322 1144 1067 148

Length:
Matches:
Conservative:
Mismatches:
Indels:

1.61e-09 601.00 31.77% 21.76% 4.55%

> Best Local Similarity: Query Match: DB:

Score: Percent Similarity:

Alignment Scores: Pred. No.: Gaps:

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122163 TACCTCAAGCGCGCTCCGGCGGGGGGGGACACCAGGCGTCTGCGCGAGATCGAG 25222
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    24888 CTCGCGCCGCCGCCGCCGCCGTCCGGAGCT-CGCCGCCCCGGCTCAGGGCCGCTGGCCGC 24946
                                                                                                                                                                                       -----ccigecaciácesecaceaciásececiciticaeacacesea 25102
                                                                                                                                                                                                                                                                                                                                                                  25103 ACACGGAACGGACAGGCGAGAACGGGAGCCATGGCGAACAACGAGGACAAGCTCCGCGAC 25162
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  25223 GGACGCACGCACGAGCGGTGGCGATCGTGGGCATGGCCTGCCGCCTGCCGGGGGGGTGTC 25282
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              164 Pro-----ProArgLeuSerLysGluGluLeulleGln----AsnMetAspArgValAsp 180
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                                                                                                                                                              87 GluSerHisSerTyrLeuProGluLeuGlyLysSerGluMetGluPhelleGluSerLys 106
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             290 AlaArgLysGlnTrpLysGlnLysPheCysGlnArgTyrAspGlnLeuMetGluAlaLeu 309
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       310 GluLysLysValGluArglleGluAsnAsnProArgArgArgAlaLysGluSerLysVal 329
LeuSerProGlySerIleIleGlnProGlnArgArgArgProSerLeuLeuSerGluPhe 72
                                                                                                               24947 GGCCCTGGGGGACGACGCGACGCCACCGACCTGGACGACGACGACGACGACGACCT
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                                                                               ------LeuHisLeuArgPro
                                                                                                                                                                                                                                                                                                                              ProlleGluSerLysHisArgSerLeuValGlnIleIleTyrAspGluAsnArgLysLys
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                                                                               73 GlnProGlyAsnGluArgSerGlnGlu---
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đ	17724 CTTGAAAAGTTGGAAGAAGAGGCTCTCCATGAGTACAGAGAAGACTCCAACGATAGAATC 17783	à	889 yAlaLeı
ò		qq	18754 CGAAGC
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ò		qa	. 18814 GGCTGG
셤		ò	922 erAlaA
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g	CCTGAGAAGCTGAAAGAGAAGAGAGAGCACACAGACACTCCTCATCTTCATCCAAAGAGGC	ò	939
<i>8</i> 8	618 GluGluMetGluThrAlaLysLySGlyLeuLeuGlu-HisGlyArgAsnITbSerAtall 63/ :::	qa	18934 ACATGA
2		λo	
3 A	AAGGAGGGGGTAGCAGGAAGGACTCCCGGCCAGTACGAAAAGGACTTCCTGGAGGCGGAT	අ _ධ ,	
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ò	luMetValGluGluAlaGl	q	19220 TGCCAG
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셤	GGAGAGGCACCGGGATGCGGATGGGTTGCGGGCTGCTGCGGGCTGCTGCGGGCTGCTGCGGGCTGCT	ò	1038 luAlaG
ò	742 lAsnAsnSerSerAspThrGluSerILeProSerProHisThrGluAlaAlaLySaspIn / 0.2	Ор	19340 CAGCTA
셤	18343 ATCACAGGACGAGCTCCTGCGGCATCACAGGACGAGCAGAAGCCCGCCACCA 18398	ò	1058 roProA
ò :	762 rGlyGlnAsnGlyProLysProProAlaThrLeuGlyAlaAspGlyProProProGLyPr 782	qq	19397 TCCCGC
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ò	811AlaProProSerProSerAlaProProProValValProLysGluGluLysGluGl 829	· 3 8	
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ф	AGGAGATGAAGCAAATGAAGAAGCTGAGGCACCGGTCCGG	λō	1168 sGlnG]
ò	869 ualaGluGluGlyProAlaLysGlyLysAspAlaGluAlaAlaGluAlaThrAlaGluGl 889	_	-

|||::: TCGACAGGITCTCCGIGGCTTCAAGIGGGCTTTCGGAAAACGCCAGCCAGCTCC 19219 ::: saagaattcagcgtcgaagacaagctcttcaggcagagcagagcgttcctgctgcctc 19339 yaHisProSerValLeuGluArgGlnIleGlyAlaIle---SerGlnGlyMetSer 1130 InLeuHisValProTyrSerGluHisAlaLysAlaProValGly-ProValThrMe 1150 .euProLeuPro-----MetAspProLysLeuAlaProPheSerGlyValLy 1168 GluglnLeuSerProArgGlyGlnAlaGlyProProGluSerLeuGlyValProTh 1188 AAAGAGGTCCTGCCTGCGTCCCCAGGCCTGACCAGAGCCGGCCCACTGGCGTGC 18993 ArgGluValileLysAlaSerProHisAlaProAspProSerAlaPheSerTyrA 1078 gProValleuProArgProProThrIleSerAsnProProLeuIleSerSer 1111 GlyLysSerArgSerProAlaProProAlaAspLysGluAlaPhe---AlaAlaG 1038 GlnLysLeuProGlyAspProProCysTrpThrSerGlyLeuProPheProValP 1058 OPro------GlyHisProLeuBroLeuGlyLeu-HisAspThr 1091 1002 AsnLeuGlnProGluSerAspAlaProGlnGln------ProGlySerSerP 1019 AlalleProprolleGinValThrLysValHisGluProproArgGluAspAlaA 990 lapro------GlnAspSerAspSerAlaThrCys-S 922 AlaAsnAlaSerproGlnLysProLeuAspLeuLysGlnLeuLys---GlnArgA 970 AspGluValAspGluAlaGluGlyGlyAspLysAsnArgLeuLeu------ 938 euLysAlaGluLysLysGluGlyGlySerGlyArgAlaThrThrAlaLysSerSe

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                                                                                                                                                                                                                                                                                         proliferation. TSG16 is useful for treating disorders associated with decreased expression or activity of TSG16, e.g. cancers, (auto)immune disorders, inflammation, complications of wound healing and infections (by viruses, bacteria, fungi, parasites, protozoa or helminths)
                                                                                                                                                                                                                                     The present invention relates to human tumour suppressor gene 16 (TSG) see AAH25689. The present sequence is the genomic sequence for TSG16, TSG16 was isolated from chromosome 16424.3. TSG16 suppresses cellular proliferation. TSG16 is useful for treating disorders associated with
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                                                                   Crawford
                                                                                                                                                                                                                                                                                                                                                                               Sequence 28906 BP; 6046 A; 8023 C; 8654 G; 6048 T; 0 U; 135 Other;
                                                                                                                                   nucleic acid representing the human tumor suppressor gene TS ful e.g. for diagnosis and treatment of tumors, inflammatory
                                                                 Kochetkova M,
                                                                   Kremmidiotis G,
                                 (WOME-) WOMEN'S & CHILDREN'S HOSPITAL
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99AU-00003771.
                                                                 Whitmore SA,
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29-OCT-1999;
                                                                 Callen DF,
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28906 632 304 1021 859 132 Length: Matches: Conservative: Mismatches: Indels: 1.09e-09 603.50 33.40% 22.56% Local Similarity: Alignment Scores: .. 02

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CAAAAGAGAAGGAGAAGATTTTTAAAGAAGATAAAGAAAAACTCAAAAAAGAAAAAGG--- 16302 16327 ACGAATATTGTAAAAAATCAGTTTCTGGAGAATGAAGACACCAAATTTAGCCTTTCTG 16386 ----AAGAGCGACTCTGTGGCCAAGCTCATCTTG 16517 :::|||||| -----TTTATAGGGAAGATTCTGCTTTTG 16326 16518 GAGACCGTGAAGGACAGAAGGAGAGAGGAGGCGGGACAGCCGGGCCCCGGGAGAAGCGA 16577 16387 ACGATCAGCGAGATCGGTGTTTTCTGACTTGTCCGATT------CATCCTTTGATT 16437 ------AAGCATAAA 16667 125 AlaThrGlyGlnProAlaGlySerGluAspLeuThrLysAspArgSerLeuThrGlyLys 144 145 LeuGluProValSerProProSerProProHisThrAspProGluLeuGluLeuValPro 164 165 ProArgLeuSerLysGluGluLeuIleGlnAsnMetAspArgValAspArgGluIleThr 184 205 AlaLysProProGluProGluLysProValSerProProProIleGluSerLysHisArg 224 185 MetValGluGlnGlnIleSerLySLeuLySLySLySGlnGlnGlnLeuGluGluGluAla 204 225 SerteuValGlnIleIleTyrAspGluAsnArgLysLysAlaGluAlaAlaHisArgIle 244 62 GlnArgArgArgProSerLeuLeuSerGluPheGlnProGlyAsnGluArgSerGlnGlu 81 16438 TCAAAGGGGAGGACAGCTGGGACTCGCCA-GTGACAGACTACAGGGACATG----------SerGluMetGluPheIleGluSerLysArgProArgLeuGlu---82 LeuHisLeuArgProGluSerHisSerTyrLeuProGluLeuGlyLys-----US-09-522-753-5 (1-2517) x AAH23705 (1-28906) 16635 GAGAAGAGGAAAGAGCAGACCGAA-Percent Similarity: 112 16488 16246 98 16303 Query Match: DB: 셤 ઢ ઠે g ઠે 셤 ò g ò 원 ઠ 요 ઠે a ઠે 셤 δ 名 ò

245	LeuGluGlyLeuGlyProGlnValGluLeuProLeuTyrAsnGlnProSerAspThrArg 264
265	GlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLysLysLysLeuIleLeuTyrPhe 284
285	LysargargasnHisalaargLysGlnTrpLysGlnLysPheCysGlnargTyrAspGln 304
305	LeuMetGlualaLeuGluLysLysValGluarglleGluasnasn 319
320	ProargargargalaLygGluSerLygValargGluTyrTyrGluLygGlnPheProGlu 339 :::
340	IlearglysGlnargGluLeuGlnGluArgMetGln
352	SetargValGlyGln 356 ::: TTTAAAGGGAAAAAAGGAAAAAACATAAAGACACACACAGGGAAAGAAGAA 17108
357	ArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluHisGluValSerGluIleIle 376
377	AspGlyLeuSerGluGluGluAsnLeuGluLysGlnMetArgGlnLeuAlaVallle 395
396 17229	ProProMetLeuTyrAspalaAspGlnGlnArgIleLysPheIleAsnMetAsnGlyLeu 415
416	MetalaAspProMetLysValTyrLysAspArgGlnValMetAsnMetTrpSerGluGln 435
436	GluLysGluThrPheArgGluLysPhe
446 17394	GlnHisPro
454	IleAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyrLeuThr 473
474	IysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArg 489
490	
510	GInProMetProArgSerSerGinGluGluLysAspGluLysGluLysGlu 526
527	LysGluAlaGluLysGluGluGluLysProGluValGluAsnAspLysGluAspLeuLeu 546 :::

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δλ	2057 erProValSerSerProSerLeuThrHisAspLysGlyLeuProLysHisLeuGluGluL 2077	4190
qq	S105 AGGCCTTGGCCTCCCTCC	Qy 2340 MetGlyLeuGluAlailelleA
à		Db 4130 AACGCTGGACACGCTATTATCG
qq		Qy 2357TrpGluGluSerP
ò		DD 4070 AGTIGGGCTGGAAAAGACAAA
qq	TGCATGCCCTTCCTCACCCCGGCTGGCACCTGCCGCTCCCTCAACTACCAG	Qy 2374 SerAlaSerLeuProAlaAlaM
δ	1yGluAlaAlaHisLeuProHisLeuArgProLeuProGluSerGlnProSerSerProFil	Db 4010 TCCCCCACAGTCCGCGTCAGT
qq		Qy 2383IleT
δ		Db 3950 TAGCTGAGCCGCTTCGCGGTAG
셤	CGCAACCTCATCTCCTCCCAACTCCTCTCCCCCCCCCCC	Qy 2397 ProGlyGlyGlyGlyLysAlaL
ò		Db 3890 GGTGGTGGTGGCGTT
ą		Qy 2410S
λō		Db 3830 GTCAGGACCATCTCATCCTCCT
qa	AACTAC	Qy 2423 SerGlyAspArgProProSerV
δ		Db 3770 ICCTCGTCCACACAATCTCCT
ΩD	4764 cecadecegriciogacidacacidraaricidecagaecegecacidrargecagaerrigie 4705	Qy 2443 ThrProLeuThrAsnArgValT
λŏ	2172 ro 2172	Db 3740
qq	4704 CGCATGAGCTACCAGGACCTCATGCAGGAGCACAACTATGATGTCACTGACTCACGCAAC 4645	Qy 2462 eproTyrAsnProLeulleMet
ζŏ	2173ValLeuAspLeuArgArgProProSerAspLeuTyrLeuProProProAspHisGlyA 2192	3713 GCC
qq	4644 GICTICGCCCGCGCCGCCGCCGGGACCCAICGCCAICGTCGACGAGTGCAIGGAG 4585	Oy 2482 oGlyLeuProAlaGly
ò		DD 3686 CGCAGCTCCTGCAGCTGCTGCC
셤		Qy 2498TrpAs
ò	SerValLeuGlyGlyGlyGluAspGlyIleGluProValSerProP 2	Db 3626 GCGCTCAGATCCACCGATGGA
g	4536 T-CCAAGCTGCACG	RESULT 83
'n	2228 roGluGlyMetThr	12370 AA
යි	4510 CTGCACCGGCTACGCCGTCCTCGTCGTCCTCCACAACATGAACCCCCGCGCGCG	
ò	2239 eralaValTyrProLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlyS 2259	XX DT 08-AUG-2001 (first entry)
q		XX DE Human tumour suppressor gene 1
ò	2259 erLysSerProGlyAsnThrSerGlnProProAlaPhePheSerLysLeuThrGluS 2278	
qo	4421CCAAGTACCACATCATGTCCCCCTCATGCAGCCCGCCCAGGTCAACCGGTT 4370	
ζō		KW autoimmune disorder; infection KW cellular proliferation suppress
qq	4369 CATC-AATGCCTACACCAAGGGTCTGCCCACCGCCATCAGCCTCCTCAAGGACATTT 4311	
ò		XX PN WO200132861-A1.
QQ	425	XX PD 10-MAY-2001.
ζ	2308GlnProGlyThrGluIlePheAsnMetProAlaIle 2319	XX XX 30-OCT-2000; 2000WO-AU001329.
qa	4250 ACCCAGCCTCCAGTGGCTTTACCTGCACCCCGCGGGGAGTCATGCCCCATGTATTCA 4191	YYY —

| IrrpGluAmpArgProSerSerAlaGlySer-ThrProPh 2462 etargleuglnAlaglyValMetAlaSerProProProPr 2482 TCCACTCCAGCAGCTCAGGGTCCCCACCACAGTCATC 3831 ValSerSerValHisSerGluGlyAspCysAsnArgArg 2442 rcricrici----- 3741 Lysval----- SerGlyArgPro----- 2409 MetPro----- 2382 TIGICCICCGCGCAIGGACCGCTCCTCTTGCTCGCGCCG 3951 ThrAlaAlaAspGlyArgSerAspHisThrLeuThrSer 2396 SG16; human; immune response modulator; cor; signal transduction activator; rapy; anticancer; anti-inflammatory; archivecome 16q24.3; alternative splicing; assor; ds. TyrArgSerGlnAlaValGlnGluHisAlaSerThrAsn 2339 GGTGCCTCGAGAAGATACACAAGACCCTGCAAGACAGAG 4131 ArgLysAlaLeu-----MetGlyLysTyrAspGln--- 2356 GITCCAAACAICCAIAAGCIITAITAITAITCICAAIAA 4071 ProproLeuSerAlaAsnAlaPheAsnProLeuAsnAla 2373 16, TSG16, genomic sequence. AspGluGluProLys 2503 AAACACCGCCCGGAA 3592 BP.

1187 7896 1207 7861 1227 7833	Oy 1243 eThrArgollelleGly Db 7777 GGCACGCCTGCAGGCCCGCTCAC OY 1250 pSerProSerArgleuAepArgGly Db 7717 CAGACCTCGCCGAGCTGGCGCAGG OY 1267 sValileTyrGluGlyLysLysGly Db 7659 Oy 1287 rGlnCysSerLysGluAspGlyArg Db 7647TGC	Oy 1307 oLysargThrTyraspMetMetGlu Db 7609 TCTTCC OY 1327 eGluGlyLeuMel Db 7567 TCGATGAGATCGTCCTGGCCGCGCG QY 1342PrOHisHi Db 7510 CCTTCCTGGTCACCGCACCTTCA	Oy 1356 hrGlnGlylleProArgSerTyrV : : :	Db 7323
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sgiyserProglyArgThrPheProProValHisProL 1490 ArgProLeuLysGluGlySerileThrGlnGlyThr- 1455 ---TyrAspThrGlyAlaSerThrThrGlySerLysL 1470 AGTICTACCGCACGGGGACTACTTACGCGATCAGGAC 7203 sesacages de la contra del la contra del la contra del la contra del la contra de la contra de la contra del la contra d JAlaLeuGluArgAlaCysTyrGluGluSer---- 1507 ProLeulysleulysproAlaHisGluGlyLeuValA 1416 SerIleHisGluIleProArgGluGluLeuArgHisT 1436 ------GAAGAAGGAGAGCCAG 7125 rgSerSerSerGlyProProHisGluThrAlaAlaPr 1307 ---CCAGCTGCGGCGCCGCCTGCGCGGGCCGCCC 7610 lugiyargValGlyarqAlalleSerSerAlaSerIl 1327 ||| ::: |||| CTGGAGGTCTACATCTCGGGCT 7568 etglyArgAlalleProProGluArgHisSer---- 1341 isLeuLysGluGlnHisHis-IleArgGlySerileT 1356 ValGluAlaGlnGluAspTyrLeuArgArgGluAlaL 1376 TICCGCCCGCCGCGACTACGCGGACTGGGAGGAGG 7391 ProProProProProSerArgAspLeuThrGluA 1396 lyHisValLeuSerTyrGluGlyGlyMetSerValTh 1287 laAspValLeuTyrLys------GlyThrIl 1243 ACCGACACCTCTGGCAGCACCTGCTGGCCCACAACC 7718 ly------ArgGluAspSerLeuProLysGlyHi 1267 uArgGlyThrAlaLeuGlySerValProGlyGlySe 1207 hrangValProSerAspSerAlaileThrTyrArgGl 1227 ..----GluAs 1250 cgaggaccccagcacagtacgr------| | | | ::: ACAC-----

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         The invention relates to a replication-defective recombinant porcine adenovirus (PAV) vector comprising at least one heterologous nucleotide sequence that lacks BlA function and retains Blssmall function. The replication-defective recombinant PAV vector is useful for preparing a composition for preventing or treating PAV infection in swine or other mammals, or for treating cancer or viral diseases in humans, e.g., AIDS. The present sequence represents the nucleotide sequence of porcine adenovirus type 3 (PAV-3) genome
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    New replication-defective recombinant PAV vector, useful for preparing a composition for preventing or treating PAV infection in swine or other mammals, or for treating cancer or viral diseases in humans, e.g., AIDS.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      PAV; PAV-3; replication-defective; E1A; E1Bsmall; virucide; cytostatic; vaccine; gene therapy; cancer; AIDS; gene; ds.
                                                                        GTCAGGACCATCTCCTCCTCTTCCATGGTGTTCGCTCCAGCGCCGGGGCGCCACC
                                                                                                                                                 2443 Thr ProLeuThrAsnArgValTrpGluAspArgProSerSerAlaGlySer-ThrProPh
                                                                                                                                                                                                eProTyrAsnProLeuIleMetArgLeuGlnAlaGlyValMetAlaSerProProProPr
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      ----SerGlyArgPro
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       2397 ProGlyGlyGlyGlyLysAlaLysVal--
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19-JUL-2002; 2002US-00199550.
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10970 CGGGGGGGTCACTCCCCCGGCTCTCCACGCGCCCTCAAAGTCCTCGGGCTGGAG 10911
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CGCAACCTCATCTCCTCCCCAACTCCTCTCCCCCCGCCGGA	2172 ro		2239 eralavalTyrProLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlyS 2259	2278 erasnSerAlaMetValLysSerLysLysGlnGluIleAsnLysLysLysLeuA 2295	2308GINProGlyThrGluIlePhe	2340 MetGlyLeuGlualaileileArgLysAlaLeuMetGlyLysTyrAspGln 2356	
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	Qy 1890 ValGluProSerLygProThrValLeuArgSerThrSerThrSerSer 1905 Db 5641 AGCTCACCTGGCTGAGTGCGAGAGCCCAGTGCGGAGCGCTGCCGAGCGCCACA 5582 Qy 1906 ProvalArgProAlaAlaThrPheProProAlaThrHisCysPro	Db 5521 CTTGCGGGCCCCGGCAAGCTCCGGGCCATGCCACCCACCC	1965 aArgSerGlyLeuGluProAlaSerSerProSerLysGlySerGluProArgProLeuVa 1965 aArgSerGlyLeuGluProAlaSerSerProSerLysGlySerGluProArgProLeuVa 1973 TTCACCGTCACCGAGACCA	2005 GHISAIASETPIOASPTOPROAIAPTOPROAIASETAIASETASPPROHISARGGULY	522 AACCAGACCTGGAGGACGATACCGTGGGACATGTAACCGAGCTCTGGGAGCGCATGC 2045	Db S105 AGCCTTCGCCTCCTCCTCCC	Db 5013 CTCAACGCCTGCATGCCCTCCTCACCCCGGCTGCCCCCCCTCAACTACCAG 4954 Qy 2098 lyGluAlaAlaHisLeuProHisLeuArgProLeuProGluSerGlnProSerSerSerP 2118

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11111 CACCIGNACCOCCTOCACCCCCCCCCCCCCCCCCCCCCCCCCCCCC	328by	10119	355	375 10014	392 ellatavalliebrokrome	404 9894 TGCCCGT	416 9834 0034	429 etasimetiipsetalusiustustustustus ::: ::: 9774 TGCGCGCCATCCGGCAGCTCAAGACGG	448ProLysAsnPheGlyLeu	458 eudlukrg	461LysThrValAlaGi	470T) 9534 GCAGCGCACCGCCCTGCGCGGGGCGCG	481 erLeuValargArgSerTyrArgArgA. ::::	499 IndlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGl ::::::::::	519 luLysAspGluLysGluLysGluLysG' 9370GCGGAGGCAGCAGGAG		559 spGluLysGluAlaValAlaSerLysG'	/ 579 ysGlyArglleThrArgSerMetAlaA.	
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	5911eGlnProGlnArgArgArgProSerLeuLeuSerGluPheGlnProGlyAsn	11171 CAGCAGCTCGCGGTCAGCGCGGGGGGGGGGGGGGGGGGG	97 LysSerGluMetGluPheIleGluSerLysArgProArgLeuGluLeuLeuFroAsgPro 	117 LeuLeuArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySerGluAspLeuThr	137 LyaAspArgSerLeuThrGlyLysLeu	146GlubroValSerProProSerProProHisThrAspProGlu	160 LeuGluLeuValProProArgLeuSerLysGluGluLeuIleGlnAsnMet	177AspargValAspArgGluIle	10790 GGCCTCTCGCTTGAGCTGCACCCGTGGGTGCCGCTCGGGGGGCCGCCGCCGCCCGACAGCCG 184 ThrMetValGluGlnGlnIleSexLysLeulysLysLysLysGlnGlnGlnGlnLeuGluGlu ::::::::::::::::::::::::::::::::	10730 CGCCAGGCCCTCCCCTCGCGCGGCGTCCAGCGACGGCG-AGGGCGACCGAGGC 204 AlaAlaLysProProGluProGluLysProValSerProProProIleGluSer [106/1 GCGTCGCCCCCTCCGCGCCCAGGCCCGGCCGGCGCTTCALCIGCCGCAGAACGGGAT 222LyBHisArgSerLeuValGlnIleIleTyrAspGluAsnArg	10611 GCATGGCTACATGAAATAAAAAACAGAAACAGAAACACACAC	10551 AAGACGGCGGACGCCCCTGCTGGAGCCTGGCAGGCTTCACTACCAAGATCACAAAAAAAA	10491 GATAGATCGCTACCACCCAACCAACCAGGTTCTTTTGACAGGGGCCCTCG 263 Thr-ArgGlnTyrHigGluAgnIleLysIleAsnGlnAlaMet	10431 ACTGCGCCCCCCTCACGGGGCAATCCATTTACACCGTGGAGCTCTTCCGCCCGGTGCGAA 277	10371 ACATTTGGAACCGCGCCACCAGTGGACCGGGGCCTCGTCTC	10326 CCGTCGGCATCGCCTGGATGTCCCGCTTCATCTACCGCTACCACCGCTCAAA 306 etGluAlaLeuGluLysLysValGluArglleGluAsnAsnProArgArgArgAlaLysG	10275 TG	

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339	lullearglysGlnArgGluLeuGlnGluArgMetGlnSerArgValG 355 AGGTCATCCAGCCCCACTACCAGGTGAACTGGTCGTCGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTCGTGGTG
355	yGlnArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluHisGluValSerGluI 375
375	lelleaspGlyLeuSerGluGluAsnLeuGluLysGlnMetArgGlnL 392
3	eualaValIleProProMetLeu
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429	etAsnMetTrpSerGluGlnGluLysGluThrPheArgGluLysPheMetGlnHis 447 :: ::
448	ProLysAsnPheGlyLeu
458	euGluarg
9654	469
9594	AGGAGGTACCTACGAGAGAGTTGACACGCTGCATCATCAGTGCGCTCAGCCTGCCCCTCT 9535
9534	CAGCGGCACCGCGCGCGCGGGGGCGCCTTCGAGCTGCGGCCCCGGGAGGGGGGCGCGCGGG994
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51	539
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559	spGluLysGluAlaValAlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgArgL 579 ::: GAAGAGCGCGGGTGGCCGGACCAGC
579	GluAlaAsnSerGluGluAlaIleThrProG
9280	92
599	InglnSerAlagluLeuAlaSerMetGluLeuAsnGluSerSerArgTrpThrGluGluG 619

(revised)
(first entry)

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            1075 ATTCGGCGGTTCCGTCCACATCGACCAACACGAATCGGTCCGGGTTCTCCGACTGCGCCG 4016
                                  -----ArgMetGlySer 2259
                                                        1015 AACGCAACAAAACCCCACGACGCAGCATGCACCAGGTCGGAAACACCCTCGGCGGACCTG 3956
                                                                                                                            2278 ------SerAsnSerAlaMetValLysSerLys-----LysGlnGluIle 2290
                                                                                                                                          2291 AsnLysLysLeuAsnThrHisAsnArgAsnGluProGluTyrAsnIleSerGlnProGly 2310
                                                                                                                                                                                       2311 ThrGluIlePheAsnMetProAlaIleThrGlyThrGlyLeuMetThrTyrArgSerGln 2330
                                                                                                                                                                                                                                      -GlyLeuGlu 2343
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                                                                                            3955 TCGACGCGCACCACACGTCACAACCACCAGGCGAGATCCGGCGAATCGGCCGCCAA
                                                                               2260 LysSerProGlyAsnThrSerGlnProProAlaPhePheSerLysLeuThrGlu-----
                                                                                                                                                                                                                                                                                                                  2344 Ala------IleIleArg
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The present sequence represents the complete nucleotide sequence of the genome of porcine ademovirus-3 (pAV-3). The specification also describes a defective recombinant PAV vector comprising inverted terminal repeats (ITR), packaging sequences and at least one heterologous nucleotide sequence (II), but lacking El functions. The defective vectors replicate inefficiently in cells (other than helper cells) so are unlikely to be immunogenic. Deletion of the El (and optionally other regions) increases the size of heterologous insert that can be packaged. The PAV-3 polynucleotides sequences are used to produce (recombinant or defective) vectors that can express heterologous proteins, e.g. for making live, recombinant virus or subunit vaccines, for nucleic acid immunisation or for gene therapy (e.g. of genetic diseases such as hemophilia or cystic fibrosis, cancer, or viral infections, including acquired immune deficiency syndrome), also for in vitro expression of recombinant therapeutic proteins. They are also used diagnostically to detect PAV antigens and/or nucleic acid. The vectors may be used in human or contingent and or contingent which may be used in human or contingent and contingents and continue continued continu
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           PAV-3
                                                                                                                    PAV-3; defective recombinant PAV vector; live recombinant virus; subunit vaccine; nucleic acid immunisation; gene therapy; genetic disease; hemophilia; cystic fibrosis; cancer; viral infection; acquired immune deficiency syndrome; PAV antigen; porcine pathogen; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          New nucleic acids from the genome of porcine adenovirus-3, and derived gene therapy vectors, particularly for immunization.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  antigens and/or nucleic acid. The vectors may be used in human or veterinary medicine, but particularly for expressing protective determinants of porchie pathogens. Regulatory regions may be used to control expression of heterologous genes. Antibodies raised against F polypeptides can also be used for diagnosis (to detect PAN-specific antigen). (Updated on 17-OCT-2003 to standardise OS field)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Sequence 34094 BP; 6240 A; 11070 C; 10693 G; 6091 T; 0 U; 0 Other;
Complete nucleotide sequence of the PAV-3 genome.
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30.98%
21.14%
4.59%
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22 HisSerLeuSerTyrProValGlnIleAlaArgThrHisThrAspValGlyLeuLeuGlu

Length: Matches: Conservative: Mismatches:

Indels:

Similarity:

TyrGlnHisHisSerArgAspTyrAlaSerHisLeuSerProGlySerIle-----

42

AAZ30163 standard; DNA; 34094 BP

AAZ30163

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805 aProThrProPr	818 oProProValve	::: 8837 CTG-CTGATCG	838 uGluGlyGluG 	8793 TGAGAACGCCAC	858 aGluGluProVe	8760 GCGGTCGCCGAC	877 yLysAspAlaG	8700 TCCCGACGCC-	89/ dGlyGlyselG		8583 GGCCAGAGCGA	929 a-Glu	: 8523 AGGAACCGCCC	934 ysAsnArgLeu	8463 ACCCCCACCTC	951	8403 TTCGTCGGGTG	956 erProGlnLys	8343 TICCGCAGITI	896	8283 CGCGCGTTCGC	696	8223 GAGACGCGCC	984 roProArgGlu	8163 GCCCAGCACG	998 laProPro	8103 CACCATCCCA	1014 lnProGlySe	8043 AGCCAGGCCG	1034 laPheAlaAl	7986 CCGCTGGAAG	1054 ProPhePro-	7947CCCG	1073 erAlaPheSe
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רבין יורייי בריז אקפרותן היוירייי בריז (פריז היוירייי בריז היוירייי בריז היוירייי בריז הייריי בריז הייריי בריז			ccacgregregergacaacgregarcaccccgregecaacgcccccccgcagagagagagagagagagagaga		CCAGCAACGACICCAGAICAGCAC	581 rgileThrArgSerMetAlaAsnGluAlaAsnSer	9772 GAITCGCGAGGICGCAGCGCCACGTGGCGGCGGCGGGGGGGGGG	933 INGLIMITATION FOR THE STATE OF THE STATE		9652 CACCGCGTTCGGTCACCAGGTGGCGGGCAACCGCCCCCCAACAGGCCCGTGCCGCCCG 9593				AGGCCAACTGCGGT					GluAspGluGluMetGlu			9314 GCTGCACACGTCACCACCACAGACGTGCCTCGGCCAACCGCGGCTCAGCCAACCCACCT		9254 TGGAGCAGAGCCAACGTCTGGGTCTCCGCAAGTGCGGCTGCGCCGGTGCCGGGATCC		9194 GCCTGTTCTCCGGCGCAGTAGACCACGACATCAGGAACCAGCGCGCTGGAATCAAGT	755 HisThrGluAlaAla-LysAspThr	9137 ITCICGGCTAGCGCCCAGATCCGCGCGGCGGCGGGAAICCGTIGCGAGTITGAIG	763	9077 AGAICGICITIACCGCAGGCAGIAICACCAAGCAGGGCCCAGCGACCCGGIICCGICGAC	774	9017 TGCGTCGCCACGGCTCCCAGGTCACCTCGAACAGCGCGTCGCGGGACAGCAGCCCCCAATIC	792 aprolleGlubroThrPro	8957 GĊĊĀGCTGTCCCGĀCGACGACGCTTCGTĀGCCTGĀĞĀĀĀĀĀTCGĀTCGĀĞGCGĀGĀĀCLĀC
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SCGTCGC	GluGluGluThrAl	31uGl	-GluGlı GGCATC	aGluGl - 	sSerSe ::::: GAACAA	aThrCy ATCGA	GACGAG	SerLeuLeuThrProThrGlyAsp ::: ::: :::		rttcgat	crcccc		ccAggg	ProPro] [CAGGA	LysPro!	GATGCG	AsnLeu(CCACGG	Argser ::: CGTAAT	oGlyAs	IleLysAlaSerProHisAla-ProAspPro 	Pro
oThrProProProAlaProPro	uLysGlı ::: GGAA	uGluGlyGluGluGlnIysProProAlaAla(::: TGAGAACGGCAGCCACGCGCCTTGCTCC	aGluGluProValLysSerGluCysThrGluGluAlaGluGluGlyProAlaLysGl 	YLYBARDAIAGIUAIAAIAGIUAIAThrAIAGIUGIYAIALeULYSAIAGIULYSLYSGI 	uGlyGlySerGlyArgAlaThrThrAlaLysSerSerGlyAlaProGlnAsp 	SerAspSerSerAlaThrCysSerAlaAspGluValAspGluAl	reccaac	ysasnargLeuLeuSerProArgProSerLeuLeuThrProThrGlyAsp		CGTCGGGTGCGTTCACCAGCAGCTGGATTTCGATGGC	roglnLysProLeuAspLeuLysGlnLeuLysGln 		CGCGCGTTCGGGGACCACCAGCGGTGTTTCCAGGGTGAGCTCTTCCACCCGGT		Thr.	GCCACG	lapropro	InProGlySerSerProArgGlyLysSerArgSerProAlaProProAlaAspLysGluA AGCCAGGCCGATACCGTCACTCGCCGTAATCCGATGTTTCCAGCCAGTACCGCTG	PhealaAlaGluAla-GlnLysLeuProGlyAspProProCysTrpThrSerGl: 	luvalIle	끞
aProPr GCCACC	OProValValProLysGluGluLys ::: -CTGATCGTCCTGCCGGAGCGGAA	raProPr GCCGCC	erGluCy CGCCTG	lagluAl agredec	laThrTh crGAAG	erAspSe :: ACTCATC	CGGTGG	ProArgi CACCGG		CCAGCA	Aspleu AAACTG	1	CCAGCG	AlaAla 	Alapro	GGGCCA	Pro rcggccA	ArgGly STCACAT	a-GlnLy :::::	ProArgGluVe	aProPro
coProAl ATCACG	1ProLy rcccrcc	luGlnLy	allysse sceces	lualaa] arcgo	lyarga] argccg	Se CGTAGA	TACCAC	LeuSer ::: TCGACC	1	CGTTCA	ProLeu		GGACCA		Aspala	CGATGCT	rcaccal	rSerPro ATACCC	laPheAlaAlaGluAla-(-	Phepro-ValbroPro	rTyrAl
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805	818	838	858	877	897	915	929	934	95	8403	956 8343	968	8283	96	984	i i	998	10143	103	105	107

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10072 GTGACCGCCGGTAATGCGCCCAGGAACGCGTTCGCCGCCGCATAATTGCCCT 10013
       CAACGCCTC 10915
                                                                                                                                          .0914 ACGCGGGGAAATCCCGAAGAAACCCGCATCGAACCGATCGGCGTCGGTGAGGAAGGCCCC 10855
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       266 TyrHisGluAsnIleLysIle-----AsnGlnAlaMetArgLysLysLeuIleLeuTyr 283
                                                                                                                                                                                                                                                                                                                                                                                                                          211 GluLysProValSer -- - ProProProIleGluSerLysHisArgSerLeuValGlnIle
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                                                                      SerlysLeutysLysInglnGlnGlnLeuGluGluGluAlaAlaLysProProGluPro
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       CGGATCCATCGC-
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       10944 CCGTTGCTG----
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                                                                                         This inventor describes a lover mechanic card it is an are encoded activity involved in biosynthesis of spinosyns. (I) are used (i) to activity involved in biosynthesis of spinosyns. (I) are used (i) to activity involved in biosynthesis of spinosyns. (I) are used (i) to activity involved in the biosynthesis of constants. (II): (ii) to generate a library of polyketide synthases; (iii) for adding forosamine or trimethylrhamnose to a spinosyn or polyketide aginosom, and (iv) for recombinant production of the corresponding enzymes, which are used for production of the corresponding enzymes, which are used for production of transgenic plants that express (II) derivatives, including production of transgenic plants that express (II) are also useful as markers for sequencing of the Saccharopolyspons. (I) are also useful as markers for sequencing of the Saccharopolyspons. (II) are also useful as also be used to raise specific antibodies, useful for identifying also be used to raise specific antibodies, useful for identifying controlles in a gene bank. Cells transformed with (I) may produce (II) at significantly increased levels or produce new derivatives of (II). This sequence, ORF 22, encodes an S. spinosa polyketide synthase
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11217 CAAATGCAACGCCACCAACGACGACGAACACGCAGTATCCACCGTCACCGCAGGACCCTC 11158
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------AATCCGGCCCGACGCACTCCCAGC 11113
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                                                                          a novel method nucleic acid (I) and its encoded
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Matches:
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7а; Page 284-313; 354pp; German.
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23725 GCGACACCCGGGCACCCCCCCCCCCCCCCCCCCCCCCC	XX DR WPI; 2001-267102/28. DR P-PSDB; AAB70969. XX PT New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for PT recombinant production of insecticidal spinosyns and their derivatives. XX XX
### CONTRIBUTION OF THE PERCENCE OF THE PERCEN	23656 CAGCACGAACGCCGCAGGTCCCGGTCCCTGGTCAG 23691 2313 IlePheAsnMetProAlaIleThrGlyThrGlyLeuMetThrTyrArgSerGlnAlaVal 2332 23692 CTCGTCCAGGTGCCGGGCGCCCTCGGCTTGGG

us-09-522-753-5.rng

ò	915 eraspSerSeralaThrCysSerAlaAspGluValAspGluAlaGluGlyGlyAspLysA 935	Db 19862 TG-GTCGGCGAGCCACGGCTGGGCCGACAGCGA
Q	19075 CACGTC 19080	1261
à	935 snArgleuLeuSerProArgProSerLeuLeuThrProThrGlyAspProArgAlaAsnA 955	19912 GGCGCCGTCCGCGAGCGGCACGAGCGCGCCCA
q	19081 GCGGGCGGTGGGGCCTCGGGCAGCTCCACCGGCGCCCACGGGTCTCCAG 19131	1268
ò	955 laSerProGlnLysProLeuAspLeuLysGlnLeuLysGlnArgAlaAlaAlaIleProP 975	19972 GGCGG
q		Qy 1283
δλ	leGlnValThrLysValHisGluProProArgGluAspAlaAlaPro	Db 20032 GACCGTGGTCGGCACCTCGGCCGTGCCCGCGCGCCCGCGTGCCG
qa		Oy 1284 tSerValThrGlnCysSerLysGluAspGlyArgSerSerSer
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à	1061 luVallleLysAlaSerProHisAlaProAspProSerAlaPheSerTyrAla- 1078	20331
q	19351 CCGCTGCCCGGTGAAGCTGCTCAGCGCGCACTCCAGCAGCGCCGGGTGCAGCGC 19410	1343
ò	lyLeuHisAspThrAlaArgProValLeuPro	20391 CGACG
đ	19411 GAACCGGGGGCCCTGCGCTGGTCGGTCGACTCGCCGAACACCTCGTCGCC 19470	Qy 1360 oArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGl
λο 1	1098 ArgProproThrileSerAsnProProLeuileSerSerAlaLysHisProSerVal 1117	Db 20451 CGGCGAAGTCGACGCC
음 ;	19471 GCGCCGCCACAGCCGGGICAG	
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3 8		Qy 1398 SInrelinatareu
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ò	1158 ProLys	N
qq	19605 GCGCGCGGGGCCGAGGACGCCGCGCGCGTGCACCCGGGGGCTGGTCGC 19658	Qy 1412 uGlyLeuValAlaThrValLysGluAlaGly
ò	1170 GluGlnLeuSerProArgGlyGlnAlaGlyProProGluSerLeuGlyValProThrAla 1189	Db 20643 GCATCGCGCCGGCCGGACAGCTCGCGGGCGATCAGCGCGC
qq		Oy 1429 oArgGluGluLeuArg
ò		Db 20703 GGCGGCCTCGGCGAGCGTGAGCGCGCCCCGCGACGTGGGCGC
đ		Qy 1435 sThrProGluLeuProLeuAlaProArg-ProLeuLysGlu(
ò	1210 LysGlylleProSerThrArgValProSerAspSerAlalleThrTyrArgGlySerIle 1229	CCGA
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------ThrGlySerL 1469 3CrccAgcgcGACCGGGTCGA 20882 GGCGGCCACCTCGCCTGCG 20762 ccagagcgccgcgagca 20822 GACGTCGCCGACGACGGTGA 20642 SerileThrGlnGlyllePr 1360 :::|||::|| GrcGACcrGcGGGAGTGCC 20450 ----- 20466 GTTGACGGCCGCCACGCACA 20582 :::||| staccagrecredecercsa 20330 GATCTCGGGTTCGCGGGCGG 20390 GACCGCTCGAACTCGCGCA 20522 -----Hi 1435 uGlyserileThrGlnGlyr 1455 erLeulleGlySerProGlyA 1482 AGGTCGACGACGCTGGTGGT 20211 GluAlaLysLeuLeuLysAr 1380 AGTACCGCTCGCGCTGGAA 20031 GGGCGAGGTCGACGGGCAC 20091 ||| ||||||| rcgAgcccgccrcGrcGCG 20151 1342 AspLeuThrGluAlaTyrLy 1398 LeuLysLeuLysProAlaHi 1411 CGAGAGCCGCCCGGTGAGGAC 19911 :|||:::|||||| |GCCCCGAGGCACGAGCCC 19971 roLysArgThrTyrAspMe 1313 lleproprogluarghisse 1341 uProLysGlyHis---- 1267 erīyṛGluGly----- 1282 -----GlyMe 1284 erglyproproHisGluTh 1304

> -	597 }	hrproGlnGlnSerAlaGluLeuAlaSerMetGluLeuAsnGluSerS	613
Ω	18172 (crecegregecegecegearegeceagegegergreatecacatecacagececea	18231
>-	613	erArgTrpThrGluGluGluMetGluThrAlaLysLysGlyL	627
Ω	18232 (gecéarédaceresceseceseces de contra de la contra del la contra della co	18291
`	627	euLeuGluHisGlyArgAsnTrpSerAlaIleAlaArgMetValGlySerLysThrValS	647
۵	18292 (cccgragccacacaratraccr	18317
>-	647	erGlnCysLysAsnPheTyrPheAsnTyrLysLysArgGlnAsnLeuAspGluIleLeuG	667
۵	18318	IČC	18321
>-	[667]	InGlnHisLysLeuLysMetGluLysGluArgAsnAlaArgArgLysLysLysLysAlaP	687
Ω	18322	geccaagccgagcaccaaggcgacgaggagagagaggggagga	18381
>-	687	roAlaAlaSerGluGluAlaAlaPheProPro-ValValGluAspGluGluMetGlu	902
۵	18382	ccroarcadcrcarcadagagagagagagagagagagagagagagagagagaga	18441
>-	107	AlaSerGlyValSerGlyAsnGluGluGluMetValGluGluAlaGluAlaLeuHisAla	726
۵	18442 (sciencesesencas con contra cont	18501
>-	727	SerGlyAsnGluValProArgGlyGluCysSerGlyPro-AlaThrValAsnAs	744
۵	18502 (seceditasasacacacacacasasasasasasasasasasasasa	18561
>-	744 I	nSerSerAspThrGluSerIleProSerProHisThrGluAlaAlaLysAspThrGlyGl	764
۵	18562 (gecsagescerceaccredececeeceaccastrogedaaccasseeceece	18621
>-	764 1	nAsnGlyProLysProProAlaThrLeuGlyAlaAspGlyProProProGlyProProTh	784
Ω	18622 (CTCCGGGCGCCCC	18636
>-	784	rProProArgArgThrSerArgAlaProIleGluProThrProAlaSerGluAl	802
۵	18637 (cccaagcagcagcagcagggggggggggggggggggggg	18696
>-	802	aThrGlyAlaProThrProPro-AlaProProSerProSerAlaPro	818
Ω	18697	GCGAGGCGCGGCCGCCGGCGGTGAGCACCACCGTCTTCGCCGGGTCCAGCCCGGCGGG	18756
>-	819	ProProValValProLysGluGluLysGluGluGluThrAlaAlaAlaProP	836
۵	18757 (CCCGGCGACCACCGGGAC	18774
>-	836 1	rovalGluGluGluGluGluGlnLysPro	851
Ω	18775	caccarcaaccaaaacaacaacaccaccacacaaarcacaaccaacaa	18834
>	851	euAlaValAspThrGlyLysAlaGluGluProValLysSerGluCysThrG	898
Ω	18835 (GGCCAGCACCCGGTCCAGCACCGCGCCCGCTCGGGGGGGG	18894
>-	898	luGlublaGluGluGlyProAlaLysGlyLysAspAlaGlubla	882
Ω	18895	SACGATCCGGCCCCGGGCTCCTCGAGCTGGGCAGAGCGCACCAGCCCCACACCGCCCCGCC	18954
>-	883	AlaGluAlaThrAlaGluGlyAlaLeuLysAlaGluL	895
۵	18955 (GACCGCCGTGCCGATCGCCGCGCGGGTGCCCGGTGACGACGAGCGCCGCCCGTCCCC	19014
>-	895	895 ysLysGluGlyGlySerGlyArgAlaThrThrAlaLysSerSerGlyAlaProGlnAspS	915
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                                                                                                  7774 AICCCGGICGAACGACTIGGGAA---CITICTICAGTIGCTGGTICAICGAACGCACCAG 7718
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Novel maytansinoid produced by bacterial host cell transformed with expression vector comprising open reading frame from ansamitocin gene cluster I of Actinosynnema pretiosum, useful as antitumor agent.
                                                                                                                                               7834 resrcescocescosrcecésrcesiècocródicéscrecciescosescréésere
                                                                                                                                                                                                           2464 yrAsnProLeuIleMetArgLeuGlnAlaGlyValMetAlaSerPro---ProProG
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                                                                                                                                                                   ----ValjrpGluAspArgProSerSerAlaGlySerThrProPheProT
                         recescorccacreccaccaccedestrarcaecerecedecaestaceedere
                                               ----LysAlaLysSerProAlaP
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                                                                                                                            -----GluGlyAspCysAsnArgArgThrProLeuThrAsnArg-
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                                                                                                                                                                                                                                                    lyLeuProAla--GlySerGlyPro---LeuAlaGlyProHisHisAla 2497
                                                                                                                                                                                                                                                              7669 GCTAGCCTTCCCGAAATCGGGTCCGGTGTTACCGGGCCCGCCTCAGCA 7621
        --GlyLysAlaLysValSerGl
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                                                 -ProSerSerArg-
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 rAspProGluLeuGluLeuValProProArgLeuSerLysGluGluLeuIleGlnAsnMe 176
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::: DESCRIPTION	1541 uThrīyrGluAsp	—გე _	1560 ValThrMetArgGluP :: :: 10750 ACTACTGTTCGACG	1580 SerGlnAspArgLysL	10690 GGCCAG	1600 ValProGluHisHisF	1620 GlyVal	 10612 CGTGTCCTGACCAAAC	1637 IleProArgGlyIle	10552 GTGTTGGCCGGTGCGT	CTGACGA	1670 eArgGly	 10432 CACCGGGTCAATTGG	1687 snAspTyrIleT	10372 ATGATCTTCGTGTTC 1706 rd3 aAsDMetLenA		1726 laGlyProArgGlyI	10301CGTGGTG	1734	10249 ATGTGTT	-		10130 GCTGTTTGTTGCCG	1775 erProLeuSerProG	10074	1793 erSerGluArgGluA ::: Canada Confedent	18081		. 1817SerThrThr	:
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Db 77501 ACCCCAAACTTTACG Qy 2300 -AsnGluProGluTy Db 77561 CAAACGGCCAGACCT		do y

AA199682 from base 600001 (Mycobacterium tuberculosis strain H. fragments LOCUS AA199682 Accession Aai99682 Begin End GCCGATATCAGCCTGAGCAGCGAATTCTGCTGACCCGCGACCG 78115 -----cgrcdacdrcaaccr------cdd 77995 ::::::
SCIBCITICGGCTTCGACACACGGTGGTCGAGTGCCGCGA 78055 CCTATGTTCGAGGCCCTCGACATCGAGCGCG------ 77937 -----GCATTCCGCATACCGAGGTGGA 77830 AACGGCGATCCCGCGCACTTTCCTACCGGCCGGTCGCCGGCGACCG 77890 GCGAACGAACTATCCACAGTGCACCCTCGATTTCCGTCGACACGTG 77560 GCGAACTCCGTTGCCCTCCCGCGACCTGCGGGAGAGATAAGGAAT 77680 GTGCGCCGGCCGTTCCGCAGCCATCAGACGGTCAAAGATG---TGCT 77797 SerProProProGlyLeuProAlaGlySerGly------ 2489 crcaacgargcrccgcggcgarattgcaggrcfrcaagccggr 77620 uGluSerProProLeuSerAlaAsnAlaPheAsnProLeuAsnAlaS 2374 oAlaAlaMetProlleThrAlaAlaAspGlyArgSerAspHisThrL 2394 yGlyGlyGlyLysAlaLysValSerGlyArgProSerSerArgLysA 2414 aproglyLeuAla-SerGlyAspArgProProSerValSerSerVal 2433 NyserThrProPheProTyrAsnProLeuIleMetArgLeuGlnAla 2473 -----ArgSerGlnAla------ValGlnGl 2334 -AsnMetGlyLeuGluAlallejleArgLysAlaLeuMetGlyLysT 2354 spCysAsnArgArgThrProLeuThrAsnArgValTrpGluAspArg [yrAsn1]eSerGlnProGlyThr-----Glu1]ePheAsn----31yPro---HisHisAlaTrpAspGluGluProLysPro 2504 111000 210000 410000 510000 710000 810000 1010000 1110000 1110000 1110000 1110000 1110000 1110000 1110000 1110000 100001 200001 300001 400001 500001 700001 800001 900001 100001 200001 300001 400001 500001 600001

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		770 ProAlaThrLeuGlyAlaAspGlyProProGlyProProThrProProArgArgThr 789	AGGTGTGCGGTGCCGCCGGTGCC AlaProThrPro AATAAACCGCGGGGGTCGGCCCC	### ##################################	73830 GGGGCCAAAGGCGGCCCCCTGCGCGCGCCACCGCCGCCCCCCCTCCTCCGCCGCCCCCCCC
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1. O. C	AA199683_07	800001	910000		QQ	72381 GATTTCGCGATTGCCACCGGTTCTCAGGTGATCGCGTCCCGAGCCGGTGTCGGAGTACGTA 72440
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3 32 3 7 07 0	AA199683_11 AA199683_12 AA199683_12	1200001	1210000 1310000 1410000		qq	12441 CTGGCGGTGGCCGGCAAGGTCATCACCCCGGGAACCCGTAACTCCGCGGGCTTCGCCG 72498
2 2 3 0 0 0	AA199683_14 AA199683_14 AA199683_15	1400001	1410000 1510000 1610000		ò	488 ArgArgGlyLysSerGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnG
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3		2111	77747		Š	680 ArgArgLysLysLysLysAlaProAlaAlaAlaSerGluGluAlaAlaPheProProVal 699

62133 CGCCAGATCAAAGACCCCCGCCACATGCGCGCCGCGATCTCACCGATCGAATGCCCGAG 62074	62073 CACCACATCCGGCCGC	62032CCAACCCCACCTGCAAACAACCCCGCCTGCGCCCACTGT	2243 roLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlySerLysSerProG	61984		y 2283 allysSerLysGlnGlulleAsnLysLysLeuAsnThrHisAsnArgAsnGluProG 2303	2303 luTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetProAlaIleThrGly	61878	61830 ACTGCCCTGACCACTGACACCACACACACACACGCTCGCAGCAACCCGTCGCGTCGCAGCAACCCGTCGCACACGCTCGCAGCAACCCGGTCGCACACGCTCGCAGCAACCCGGTCGACACGCTCGCAGAACCCGGTCGACACGCTCGCACACACA	2336 laSerThrAsnMetGlyLeuGluAlaileileArgLySAlaLeuMetGlyL	61770	61710		61650 CAGCGACCAGCCCACALCLISCACCGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGAG	61593	2409 roserSerArigiysAlalysSer	2421	61473 CAGCCCATCGGGCACAGACTCCA	2425 -AspargPro	2439	61353 CAACCGCACCGCAATCCCACTCCACATGCGGCACGCACACCGCACGCA		61293 CGCCGGCACCACACCA	61254	2487GlySerGlyProLeuAlaGlyProHisHisAlaTrpAspGluGluEroLysP	Db 61194 CAACCACAACGGACGACGGCCCCCTGCCCATACGTGGCCAGCAGGCCTGCGC 61135
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63040 CCGCCACCAAAACGCACCCGCCACCACATGATCAG-	1890 ValGlubroSerLysProThrValLeuArgSerThrSerThrS 	1910 AlaAlaThrPheProProAlaThrHisC	62950 GGTCCGCGAACTCCACCGCCG	1930 ProThrLeuMetGluProValLeuLeuE	1950 ArgProArg	62875 CCCCAGAACGGCCGCTCAACCAGAACCGCTCCCGCTGGA 1964 ProAlaArqSerGlyLeuGluProAlaSerSerProSer	62815 CCAGCCCAGGCACACGATCACCCGCGAA	1977LysGlySerGluProArgProLeuValPro- 1977LysGlySerGluProArgProLeuValPro- 1977	1987ProValSerGlyHis	62695 AGGACACCAGTACCGGTTCGGGGCCAT	2001AsnLeuAlaProHisl	2017 SerAlaSerAspProHisArgGluLys'	62576 ACTIGGGCGATGCCGGGCTGGAAGAGCACTGTCTGGCGTACATGGCGTGCCCAG		2057 SerProValSerSerProSerLeuThrHisAspLysG	2075 GludluleuAsplysserHisleuGlu	62451 CGTGAACTTGACCTCTCGTATCGCTTC	2095 LysLeuGlyGlyGluhlahlaHisLeu	62391 CGAGTGGAAGGCATGACTGACGCICAG	62331 CCCAGCAATCCGATCCACCTCACC	2129 sGlnArgValValThrLeuAlaGlnHi	62283 ATCAGGTGTTGACCGCCCCALALICACACCACACACACACACACACACACACACACACA	2149 IGHIBELEFICOLOGICA CONTROLLE CO	2167 roGlyAlaSerCysProValLeuAspLeuArgArgProProSerAspLeuTyrLeu	62187 CTCAGGCACCCACCCATCAAACGGGCCCGCGCCCCCAACCACCAGGAGCAI-	2187 roProAspHisGlyAlaProAlaArg(
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ò	159 GlubeuGlubeuValProProArdLeuSerLysGluGlu 171	λ	434 uGlnGl
; <u>8</u>	::: CAGCTCCCTCCAGCAAGCCGCACACCTCATCAAACG	q	67821 GGGCGT
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3 8		qa	67761 CGGCGG
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g (CICACCAGCGGCAGCCGGCCCGGCCGGCAGCTCAGCCCCTCTCACGCCCCGCCGCCCGC	ò	513 oArgSe
કે 1	ProprolleGluserLysHlsArgserLeuValGlnleLleTyrAspGluAsn	Dp	67619GC
gg	CACCGCCCGCTGCTCGAACAGCGCACGCACGCAGCAAAGACCAGCCCACGTCGGTGAG	ò	533 uGluLy
ò	ArgiysiysAlaGluAlaAlaHisArgileLeuGluGlyLeuGlyProGlnValGluLeu :::	q	67580
q G	CGGCGCTCGGGACGCGTCACCGGCAAAGTCGCGCAGAC-GCAGGGCCTGAT	ò	553 rSerG
ò	255 ProLeuTyrAsnGlnProSerAspThrArgGlnTyrHisGluAsn1leLys1leAsnGln 274	q	::: :::
ପ୍	68463CACGCAGGCCTGCGGACTACGGGCCGACACCACGGCACCACGAGGCCGT 68410	}	195
ò	275 AlaMetArgLysLysLeulleLeuTyrPheLysArgArgAsnHisAlaArgLys 292	i	200000 0575
ą	68409 CGGAAGCGCCGGGGGTTGCGGCCCTTCCGGGGCGTCGTCGTCGGCGG	3 8	20012000
ò	293 301 GlnTrpLysGlnLysPheCysGlnArg	Š	564 IAIASE
qa	68349 GGGCCTGTTCCACGATCAGGTGGGCGTTGGTGCCGCTGATGCCGAAGGAGAGACCC 68293	අු	67410 CCTTGP
ě		<i>&</i> .	582 leThr
े द		q	67350 GCACCC
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ò	321ArgArgAlaLy 325	7 1	
Ωp	68172 CGCCGTGGCGCATCGCCATCTTGATGACGCCTGCGACGCCCGCAGCCGCTGCG 68113	g ,	-77199 GG1CC-
ò	325 sGluSerLysValArgGluTyrTyrGluLysGlnPhePro 338	Š	
q	68112 TGTGGCCGATGTTCGACTTCACAGAGCCACAGCGGCCGGTCGGCGGGGCGGGC	qq	67212 AGAGGC
è	-GlulleArd[waG]nArdG]uLeuG]nG]uArdWetG]nSerArdVa]G]vG]nArdG	ò	651 snPheT
i		qq	67153
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q n	67941 GGGCCTGGCGGATCACGCGJJTCCTGGGCGACGTCGTTCGGGGCGGTCAGGC 67891	ò	706 GluAla
ò	394 lileProProMetLeuTyrAspAlaAspGlnGlnArgIleLysPheIleAsnMetAsnGl 414 :::	. qq	67011 CTCGC1
qq	67890 CGTTGCTCGCGC 67879	8	725 sAlaSe
ò	414 yLeuMetAlaAspProMetLysValTyrLysAspArgGlnValMetAsnMetTrpSerG1 434	: A	
q	67878 CGTCCTGGTTCACGGCCGATCCGCGTACGACGCCGAGTACTTCATGTCCGTTGCGGC 67822	ìè	
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BProLygAsnPheGlyLeuIl 454 :: ccrcGGAGAATCCGGTGCCGT 67762	uTyrTyrTeuTh 473	493	6.3	nGlnGlnProMetPr 513 ::: GGCGCAGGCCTG 67620	uAlaGluLysGluGl 533	67581	uLysThrAspAspTh 553	AGGTGTAGGCGATGC 67531	095	CGACCTCCGGCGGA 67471	LysGluAlaVa 564	ceccegréadegrér 67411	582	6	hrProGlnGlnSerA 602 ::::: IGCCGAAGAAGTCGG 67291	T 615	AGGTCGTGCCGGGGT 67231	isGlyArgAsnTrp- 634	dedregegredr 67213	erLysThrValSerGlnCysLysA 651	1.671		LysLysLysAla 686	CGATGGCGATCGGTT 67072	GluAspGluGluMet 705	ceacatrceceerca 67012	725	CGTTCGTACCCGGCA 66952	e 745	CTCGAAGAGCTCGTC 66892	
uLysPheMetGlnHisProLys 	nrValAlaGluCysValLev AGCGGCCGTCCGGTGC	erLeuValArgArgSerTyrArg	-GAGTCCGCGCTGGCGTGAGAACTCCACGAACGCCGTCGGCGTGG	nglnglnglnglnglnglr accccadecccagrodccr	pGluLysGluLysGluLysGlı	ACCAGCGATGCCGAGCACGC	uLysProGluValGluAsnAspLysGluAspLeuLeuLysGluLysThrAspAspTh	accecedecerceagecea		ccedadaridacerrocadadegrrcccorrocraeceraecercoacerccoscosoa		rggctggacacccccgcgtacac	laAsnSerGlnGlyArç	ייריפרוריפים בייים ב	AlaAsnSerGluGluAlaIleTk :::::: 	JeudanGluSerSerArg	 	AlaLysLysGlyLeuLeuGluHisGlyArgAsnTrp		aArgMetValGlySerLysThrVa			ArgAsnAla-ArgArgLysLysLysAl	GGTAGCGCAGCCATCGCGA	-SerGluGluAlaAlaPheProProValValGluAspGluGluMet	secaectereacecercretc	nGluGluGluMetVal-GluGlı	TTGTCCTCGGTACCCGGCATGT	rgGlyGluCysSerGlyProAle	AAGGCAGTTCTCGGTCGATCAG	
uGlnGluLysGluThrPheArgGluLysPheMetGlnHisProLysAsnPheGlyLeuIl 	eAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyrTyrLeuTh : :GGGGGCCTCGGCGACGGCTTGCAGCGGCCGTCCGGTGC	rLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArgArgGlyLysS	GAGTCCG	rginginginginginginginginginginginginging	oArgSerSerGlnGluGluLysAg		uGluLysProGluValGluAsnA6	CGTGTCGAGCGTC	rSerGlyGluAspAsnAspGlu	geccedaareacecrecadaec		ccceecrccceaacareresrccreecresacacccccccaracaceccesreada	1AlaSeriyeGlyArgLysThrAlaAenSerGlnGlyArgArg-LyeGlyArgI	C.T.TGAGCGTGGCGGGGTCGATGC	eThrargserMetAlaAsnGluAlaAsnSerGluGluAlaIleThrProGlnGlnSerA GCACCCGTTGCTGCGGGGCGCGGGGGGCGCGCGGGGTTGATGCCGAAGAAGTCGG	laGluLeuAlaSerMetGluI	 corgaageggggggggggggggggggggggggggggggggg	rpThrGluGluGluMetGluThrAl	GGTCC	SerAlaIleAlaArgMetValGlySerLysThrValSerGlnCysLys	AGAGGCCCGCCACGICCCAGCCGCGGICGG		euLysMetGluLysGlu	GCTCGGGGGGGGCACCCCGCCCGGGTAGCGGCAGGCCATCGCGACGATGGCGATTCGGTT	ProAlaAlaAlaSerGluGlu	ccectegedetrecreeaegregedaggegeraagggrengregagareg	GlualaserGlyValSerGlyAsnGluGluMetVal-GluGluAlaGluAlaLeuHi	zrcéciricadóragregegégages	sAlaSerGlyAsnGluValProArgGlyGluCysSerGlyProAlaThrValAsnAsnS 	rgtcttctccactcctgatcaag/	
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The invention relates to a DNA sequence which is a fully defined sequence of 103551 base pairs appearing as ABX04971, or its variant, that it is not a sequence encoding all or part amino acids 1-920 encoded by mon Al as given in the specification. The DNA is the S. cinnamonensis polyketide antibiotic monensin biosynthetic gene cluster. Also included are a
                                                                                                                                             6407
                                                                                                                                                                                                                                                                                                                        gArgThrProLeuThrAsnArg-----ValTrpGluAspArgProSerSerAlaGlySe 2459
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          6552 -----ccaacacaccaccaarccaarccaarccaraccaracacacacacacacccrc 6604
, DNA sequence encoding polyketide synthase, useful for the production polyketides such as antibiotic monensin.
                                                                                                                              | ::: |||||||
6348 AGCICGGCIACGGAIACGGICCGGCCTTCCAGGGCCTGACGGGGCTGTGGCGCGACGGCG
                                                                                                                                                                                                                                                               ccancacectroeccedaanccdecroeccececacacacadadececegegerrente
                                                                                                                                                                                                                                                                                                                                                                                    2459 rThrproPheProTyrAsnProLeulleMetArgLeuGlnAlaGlyValMetAlaSerPr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 P-PSDB; ABG99854, ABG99855, ABG99856, ABG99857, ABG99858, ABG99859, ABG99866, ABG99866, ABG99860, ABG99870, ABG99877, ABG99887, ABG998876, ABG998876, ABG998876, ABG998877, ABG998886, ABG99880, ABG99881, ABG99882, ABG99881, ABG998881, ABG998881, ABG998881, ABG998881, ABG998881, ABG998881, ABG998881, ABG998881, ABG998881, ABG998887, ABG99887, ABG99887, ABG998887, ABG998887, ABG998887, ABG998
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           2479 oProProProGlyLeuProAlaGlySerGly------
                                                                                                                                                                                                         -----AspArgProProSerValSerSerValHisSerGluGlyAspCysAsnAr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         Monensin; gene; cluster; polyketide synthase; antibiotic; ds; antihelminthic; insecticide; immunosuppressant; antifungal; antibacterial; polyether; mon BI; mon BII; mon CI; mon CII; mon H; mon RI; mon RII; mon T; mon AX.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       S. cinnamonensis monensin type I polyketide synthase gene cluster.
                                                                                     Y------beuAlaSerGly-----
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     6605 CACGCCGAAGGGGCCACCGCCTCCGCGTACGGATCACACCCA 6647
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Claim 1; Page 116-195; 212pp; English.
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cc recombinant cloning or expression vector comprising the gene cluster, a transformed host cell which has been transformed to contain the gene cluster (and is capable of expressing a corresponding polypeptide), a hybridization probe derived from the gene cluster (for identification and isolation of the same or analogous gene cluster, e.g. one which binds control expression of the monensin gene cluster selected from mon to specifically to a region of the monensin gene cluster selected from mon ax b, the use of the mon IX men en variant and a monensin prometr to control expression of a heterologous gene in Streptomyces cinnamonensis, control expression of a heterologous gene in Streptomyces cinnamonensis, control expression of a heterologous gene in Streptomyces cinnamonensis, control expression of a heterologous gene in Streptomyces cinnamonensis, control expression of a variants), an epoxidase enzyme encoded by mon CII, producing s. cinnamonensis capable of cyclase enzyme encoded by mon CII, producing s. cinnamonensis to production of monensin comprising engineering it to cyclase enzyme encoded by mon CII, producing s. cinnamonensis copies or express the mon RI gene, S. cinnamonensis containing multiple copies or express the mon RI gene, S. cinnamonensis with DNA convexpress the mon RI gene, S. cinnamonensis with DNA convexpress the mon RI gene and/or its variants, expressing agene under control of the activator gene mon RI or actil/orf4 and 13-propyl expromycin A. The convexpress and materials (enzyme expressing the gene under control of the activator gene mon RI or actil/orf4 and 13-propyl expromycin A. The cueful for preparing polyketides by recombinant synthesis. The cueful as insecticides, antiblotics, antiblotics, antiblotics, antiblotics of polyketide are useful as insecticides, antiblotics, antiblotic polyketide. The present sequence represents the monensin gene cluster gene is useful for the production of monensin, an antibiotic polyketide are useful for the production of monensin, an antibiotic polyk

Alignment Scores:

Alignment Scores:

1.33e-09
Matches:

Conservative:

Best Local Similarity:

1.5\$
Matches:

Conservative:

1.25\$
Matches:

1.25\$

US-09-522-753-5 (1-2517) x ABX04971 (1-103599)

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	11 28 :: FC 69207	44 CT 69147	53 3C 69093	ro 66	cg 69033	82 GA 68973	er 98 CC 68916	eu 118 :: TC 68871	sp 138 68859	ro 158
1	STIYTETON CAATCCGA	yrglnHis- ::: sccccaca	GCCTGCACC	rgArgArgP 	cccaccac	eu TCACCGATC	euGlyLysS ::: cGGGCCAAC	spProLeul :::: ACCGTGTGA	euThrLys?	isThraspE ::: :GAACCAACC
	9 AlaGlnThrTrpArgalaThrGluProArgTyrProProHisSerLeusSeriyrFroval 28	29 GlnIleAlaArgThrHisThrAspValGlyLeuLeuGluTyrGlnHis 44	45 HisserArgAspTyrAlaSerHisLeu53 45 HisserArgAspTyrAlaSerHisLeu53 69146 PARCAGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGG	54SerproGlySerllelleGlnBroGlnArgArgPro 66	3 - 1 1 3 3 3 3 3 3 3 3	67 SerleuLeuSerGluPheGlnProGlyAsnGluArgSerGlnGluLeu 82 ::: :: :: ::: ::: 69032 ACACGCCACATCACACACACACACACACACACACACACAC	83HisLeuArgProGluSerHisSerTyrLeuProGluLeuGlyLysSer 98	99 GluMetGluPheIleGluSerLysArgProArgLeuGluLeuLeuProAspProLeuLeu 118	119 ArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySerGluAspLeuThrLysAsp 138 :::	139 ArgSerLeuThrGlyLysLeuGluProValSerProProSerProProHisThrAspPro 158
	yrProProH TCCCGCCAC	spvalGlyL GAGTCAGGT	GCCAGCTCG	erllelleG	: :ATCAAACGC	AsnGluArgS	isserTyrI GACCGAC	Argleuglul :GCCTGCGC	ProAlaGly	SerProPro(
	luProArgT CCCACGC	hrHisThrA ATCACGTC	isLeu ::: ACATCGCG	erProGlyS	PACCCACC	ProGlyA	rogluserF - cgcAcccc-	ysargPro/ ::: \aacaaccc	rhrGlyGlnf	3luProVal
	rgalaThrG STCTTACG-	TT	yrAlaSerH	88	CCCTCAGGC	luPheGln- :::::: :AGATCAAAG	isLeuArgF ::: :ACATCCGGC	leGluSerI ::::: TGCAACGC	euLeuAla7	ArgSerLeuThrGlyLysLeuG ::: cgGGCCCAGAACACCACTC-
	lnThrTrpA accccTTC	lealaarg- :: mcacccgac	erArgAspT		ATCGCCCCA	euLeuSerG :::: :GCATCCGCC	ccgagcacc	letGluPhel	119 ArgProSerProLeuLe::	SerLeuThr(::: GCCCAGAA(
,	Alag GCTC	GlnI	Hiss -		ACAC	SerL ::: ACAC	ATGC	Glul CAC-	ArgE ::: CAAC	Args CGGC
1	69260	29	45	54	69092	69032	83	99	119	139

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3278 CGCGAGGGGGATTCCTGCACGACGCCGCCT		3338 CCCCGCGAGGCCCTCGGCATGGACCCGCAGCA	GCCGTGGAGCGCGCAGGGCTCGAC	1601 roGluHisHisProHisProIleSerPro	3458 GTCGGCGCCCCCTGGACTACGGCCCGCG	1620 lyValAspLeuTyrArgSerHisIleProLeu	3518 GGCCACCTCCTGACCGGGACCACGCCCAGCGT	GGCCTCA	1656 laproAsnProThrTyrProHisLeuTyrPro	3638 CACCT	1676 hrAlaAlaLeuGluAsnArgGlnThr 			3716 GGCATGTTCGTCGAGTTCTCGCGGCAGCGCGCGT		1718 erSerLeuAlaLeuAsnTyrAlaAlaGlyPrc	 	1734	3896 GTCAACCAGGACGCCGCCTCCAACGGGCTCAC	1746	3955 CGTCATCCGACAGGCCCTGGCCGACGCCGGGG	1756	4015	1776	4075	1792	4135 deccacecededeccecedecederade	1809 GluArgGluLysSerIleLeuThrSerThrT	4195 CCACGGCGTCCTGCC	1828	4230 GGCCCACCACCGCGTCGACTGC	1840 Ser	000
	arddlyhrddluhabSerLeuProLysGlyHisVallleTy 1270	2358 CCGIACGCTTCCTCGACGCCGTACGCACCTGGAGGATCGGGCGCCGACGACCTTCCTGG 2417	sValLeuSerTyrGluGlyGlyMetSerValThrGlnCysSe 1290		3SerSerSerGlyProProHisGluThrAlaAlaProLysAr	CCCTGCGCAAGGCCCGCCCGGAGCCCCAGTCGCTGCTCGCTG 253/	2562			1349 sHisIleArgGlySerIleThrGlnGlyIleProArgSerTyrValGluAlaGluAs 1369 ::: :::	ACGGTCAGGGTGCCCC-TGCCGACCTACCAGGGGAACGCCACTGGTTCGAC 2659 Qy		1408	כיו	1428	2780 GGTGGCGGTGATCGGCCGCGCGCCCCACGAGACGACCGAGCGCGTGCGCGCACGCTC 2836 Qy	1448		leThrGlnGlyThrProLeuLysTyrAspThrGlyAlaSerThrThrGlyS		AspValArgSerLeuIleGlySerProGlyArgThrPheProProValH		1488 isProLeuAspValMetAlaAspAlaArgAlaLeuGLUArgAlaCySiyrGlUGLUSELD 1909	CGCCCTCGCCCCCACCTGGGCGACCTGCTCACCGGCGGGCG	1508 euLysSerArgProGlyThrAlaSerSerSerSerGlyGlySerIleAlaArg- 1524 Db		alileValProGlubeuGlyLysProArgG	TACCCCGGCGCGTCACCTCCCCC	1538 InSerProLeuThrTyrGluAspHisGlyAlaProPheAlaGlyHisLeuP 1555		1555 roargGlyserProValThrMetArgGluProThrProArgLeuGlMGlu 1571 Db	OY GlySerLeuSerSerEysAlaSerGlnAspArgLysLeu 1585 Db	
			Oy 1270	Db 2418 A		2478	Db 2538 (Db 2563			Qy 1369			Qy 1408	Db 2780	Qy 1428	Db 2837		Db 2891								Db 3104		3158	Oy 1555		ı

| InpropheserSerArgHisSerSerSerP 1776 Arg------GluArgAspArgAspArg 1808 GlyThrGluGlnSerSerGlySerSerGly 1839 GCGGCGTCGAGCTCCTCACCGAGGAGCGGG 4289 -----SerGlyGlyGlyGlyGlySerSer 1848 hrLysProThrThr------ThṛṢ 1792 ThrThrValGluHisAlaProlleTrp--- 1827 ::: |||||| -----CAGGACGCTCCACG-TGGACC 4229 GCCGGCGGACCGTCATGTCGACACCG 3715 -----SerProArgGlus 1718 roArgGlyIleIleAsp------ 1733 ACCGCCC-CCAACGCCCCGTCCCAGCAGCG 3954 |yThrPro-----AlaThrAlaMetA 1756 GCTCCAGCTCGCTCGAAGTCGAACAT 4134 TyrGluHisLeuLeuArgGlyValSerg 1620 eualaPheAspProThrSerIlePro--- 1638 | | | | :::||| stgatgtcgcctaccagctc 3577 aAlaAlaTyrTyrLeuProArgHisLeuA 1656 ACGGCCTGCTCGTCCTCGTCGCGCTG 3637 roProTyrLeulleArgGlyTyrProAspT 1676-----ddccgrcctcdTrcc 3655 nrileileAsnAsp---TyrileThrSerG 1693 GCCGAGGGCGTCGTCCTCGTCGTCGAG 3835 cccdraccccrahrccaaccacccc 3895 roHisLeuProValLeuValPro----- 1745 ::: GCCTCAAGGCCAGCCGGCCGTCTTC 3457 ceraceaceacececeaededecereae 3517 TGTTCGACGCCGGCTTCTTCGGGATATCG 3337 AGCGGCTGCTCCTGGAGACGGCATGGGAG 3397 ulleAlaLysSerProHisSerThrValP 1601

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571 ThralaAsnSerGlnGlyArgArgLysGlyArglleThrArgSerMetAlaAsnGluAla 590
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/product= "vep ORF 1 amino acid sequence #1 (AAY77177)"
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/product= "vep ORF 1 amino acid sequence #3 (AAY77199)"
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The invention relates to an isolated and purified nucleic acid segment comprising a desosamine biosynthetic gene cluster, a fragment or its comprising a desosamine biosynthetic gene cluster, a fragment or its biologically active variant, where the nucleic acid sequence is not derived from the eryC gene cluster of acid sequence is not controlled sequences antibioticus. The invention also relates to a macrolide biosynthetic gene cluster encodes proteins which synthesise methymycin, parabomycin, narobomycin or a combination of these compounds. Recombinant or augmented cells comprising the desosamine compounds. Recombinant or augmented cells comprising the desosamine compounds. Recombinant or augmented cells comprising the desosamine of biologically active macrolides. The macrolide biosynthetic gene clusters are useful for the production of polyketide synthesis mader of biologically active macrolides. The macrolide biosynthetic proteins of are useful for synthesis of methymycin, pikromycin, neomethymycin and creating to prepare novel antibiotics and polyhydroxyalkanoate (PHA) carbomycin. The alternative termination of polyketide synthesis may be useful to prepare novel antibiotics and polyhydroxyalkanoate (PHA) compounds produced by the recombinant host cells are useful componers. The compounds produced by the recombinant host cells are useful as biopolymers, e.g., in packaging or biomedical applications, to componer such as chemotherapeutics, immunosuppressants, agents to treat asthma, chronic obstructive pulmonary disease as well as other diseases involving respiratory pathogens, as well as viral parasitor pathogens, or as crop protection agents (e.g., fungicides or insecticides) via expression of polyketides in plants. The present sequence represents a Streptomyce polyketides in plants. The present sequence represents a streptomyce or antibiotic which are active against or insecticides) via expression of polyketides in plants. The present sequence represents a streptomyce or antibiotic which are not sequence ins
                                      #2 (AAY77178)"
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                                      /product= "vep ORF 1 amino acid sequence
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15872 520 214 964 670

Length: Matches: Conservative: Mismatches:

3.15e-10 614.50 31.11% 22.04% 4.65%

Indels: Gaps:

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1175 gGlyGlobal 1178 1178 gGly	CGATGCTCCACGACTCCGCCGGGGTCGTCGAAGAGCTGCACCTCCAGGAGCCCCGGGGTCGArgval ProSerAspSerAlaIleThrTyTArgGlySerIleThrHisGlyThrPr	2358 CCGTACGCTTCCTCGACGCCGTACGCACCCTGGAGGAATCGGGGCCGACACACTTCCTGGA 2*17 1270 rGluGlyLysLysGlyHisValLeuSerTyrGluGlyGlyMetSerValThrGlnCysSe 1290 1271 rGluGlyLysLysGlyHisValLeuSerTyrGluGlyGlyMetSerValThrGlnCysSe 1290 1272 rCygCcGGGGGGGTCTGCTCGGGAGGGGGGGGGACTCGGTACGGGAGG 2*477 1273 rCygCacGGGGGGTTTCCGGCGATGGGGGGGGGGGGGGGGGGGGGG		
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1092 TCATCAAGACGCTCCGGGCTCGGCCGGGAGCCTCAACTTCCGTA 1151	884 GlualaThralaGluGlyAlaLeuLysAlaGluLysLysGluGlyGlySerGlyArgAla 903 1272 CCAACGCCACGTCGTCCTCAGCGAAGGCCCGCCCAGGGCGGCGGCGAGCAGCCGGCATCG 1331 904 ThrThralaLysSerSerGlyAlaProGlnAspSerAspSerSerAlaThrCysSerAla 923	AspGluValAspGluAlaGluGlyGlyAspLy8AsnArgLeuLeuSerProArgProSer	14/3	1022 LysSer	AlaProAspProSerAlaPheSerTyrAlaPro-ProGlyHisProLeuProAlaProAspProSerAlaPheSerTyrAlaPro-ProGlyHisProLeuPro	1120 gGlnIleGlyAla11eSerGlnGlyMetSerValGlnLeuHisVa 1135 1120 gGlnIleGlyAla11eSerGln
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		à	2425 pArgProPro
ò	2137 nHislleSerGluVallleThrGlnAspTyrThrArgHisHisProGlnGlnLeuSerAl 2157	q	8136 GCGGCG
셤	5008GAACAGGAACCACGCGCTCAACAGC 4983	è	2445 uThrAsnArd
ò	2157 aproLeubroAlaProLeuTyrSerPheProGlyAlaSerCysProValLeuAspLe 2176	3 A	
ą	4982 GCCGGAATCGCGTGCGGTCGTCCCGACCAGCGCCCTGTCCAATCGAGCGGCA 4923	Š	2465 nProLeulle
ò	2176 uArgAroProSerAspLeuTyrLeuProProProAspHisGlyAlaProAlaArgGl 2196	qa	: 4067 TCCGACTGGG
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č		3 4 1 1 1	AAT68715 standard;
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qq	recerrececc	岩	Streptomyces venezu
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ò		YY SO	Streptomyces venezu
ΩP	4721 GCATCCAGGAACGTGTTCGCCGCGCGTAGTTGGCCTGCCCTGCGGCACCCGCAATCCCG 4662	₹E¦	Key
ò		FF	CDS ZO
g	4661 GAGAAGGAAGAAAAAGAACGAAGAAGACGACAGATCCAGATCGCGGGTCAACTCGTGCAGA 4602	FT	
λõ	2290AsnArgAsnLysLeuAsnThrHisAsnArgAsnGluP 2302	FT	CDS 14
d G	4601 TGCCATGCCCCGTCACCTTCGGGGCAAACACCGTGCCCACATCCGCCGCCGACAAGGAC 4542	XX N	WO9722711-A1.
ò	ıTyrAsn	XX OG	26-JUN-1997.
qa	GGTATGCACG	X	18-DEC-1996; 96W
ò	2322 ThrdlyLeuMetThrTyrArgSerGlnalaValGlnGluHis 2335	×	19-DEC-1995; 95U
qq	4481 ACCGGAATGGCGGCCAGCACGTGCTCCAGATCGGTGCGGTCAGCCACATCGCAAAGCCACC 4422	XX &	(MINU) UNIV MINNE
ò	2336 AlaSerThrAsnMetGlyLeuGluAlaIleIleArgLysAlaLeuMetGlyLysTyrAsp 2355	XX	Sherman DH, Willi
q	4421 ACCTCAACCACG 4410	X H	WPI; 1997-341701/3
ò	2356 GlnTrpGluGluSerProDroLeuSerAlaAsnAlaPheAsnProLeuAsnAlaSerAla 2375	数 数	P-PSDB; AAW19629,
QD		PT	Expression cassett provide wide range
ò	2376 SerLeuPro-AlaAlaMetProlleThrAlaAlaAspGlyArgSerAspHisThrLeuTh 2395	r X	use.
ą	4361 TTCCAGCCAGCCTGCCAACACCCAAACGCCGAATCCCGTGCT 4316	8 X	Claim 54; Fig 23;
λo	2395 rSerProGlyGlyGlyGlyLy82402	88	Streptomyces venez synthase (PKS) gen
qq	4315 -CCTCCACCAGGAGGAGGGGAACCACTCCCCCAACGCACCGGTGCCACCCGTGATGAGG 4257	មួន	AAW19629-30 and AA module and a 3' en
ò		88	-ACP, an acyltrans carrier protein do
Ωp	4256 ACGGTCCCGTCTCCAGTTGCGGGGAGGACCCTCCTCGCGCACCGTGAGTCGCGCC 4197	88	hybridisation stra cassette encoding
ò			from the Streptomy monomer synthase a
qa	4196 AACCGAGGCACCACGCCGCGCGCGCGAAGTGCCAGGTGCGGTTCTCCTGCTCGCACG 4137	⊖ ×	insect) cells lor

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venezuelae vep ORFI (AAT68715) comprises the polyketide of 3, gene cluster encoding a polyene of 12 carbons (see also and AAW00918). It contains 5 PKS modules, with a 5, loading 1. transferase, a dehydratase, a keto-reductase and an acyl in domain. The gene cluster was cloned using a heterologous is strategy from a genomic DNA library. A novel expression oding the first module from the vep gene cluster and module 7 ptomyces tylP gene cluster has polyhydroxyalkanoate (PHA) isse activity and can be used for PHA prodn. in host (esp. for use as a biodegradable polymer
                                                                                                                                                                              eMetArgLeuGlnAlaGlyValMetAlaSerProProProProGlyLeuPr 2485
                                                                                                                                                                                                                         GCTGAACTCAACACCCCCACGACGACGCACCAAGGTCAGGTCAGGTCA 4008
2425 pArgProProSerValSerSerValHisSerGluGlyAspCysAsnArgArgThrProLe 2445
                                           -----GCAA¢Ġ¢ċrigc 4119
                                                                                      gvalTrpGluAspArgProSerSerAlaGlySerThrProPheProTyrAs 2465
                                                                                                                     tes for production of poly:hydroxy:alkanoate(s) -
ge of biodegradable polymers for medical or industrial
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            se; polyhydroxyalkanoate monomer synthase;
te; biodegradable polymer; vep gene;
ring; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   zuelae polyketide synthase vep ORF1.
                                                                                                                                                                                                                                                                         rGlyProLeuAlaGlyProHisHisAla 2497
                                                                                                                                                                                                                                                                                                        31.
AAW19630, AAW00918.
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20. .13912
/*tag= a
14056. .14136
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/*tag= c
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1317 gValGlyArgAla-IleSerSerAla-SerIleGluGly		7950 CTTTGCCCCGGTACCGGCGAAAAACGACGGCCAGTCCAC		7890 CTGCGCCAATGCCGTTGTCACCGTGTGCGCCTTGTCGCGCCTCGTCGCGCCTGGTGCGCCTTGTCGTGGTG		1349 HisHisIleArgGlySerIleThrGlnGlyIleProArg	7770 CACCCCATCCGGACCGACCGACGACGGTGC-TCACAC	7711CACCGTCGCGAACGGACGCTTCTCGCACCT	1387	7657 TAGCCGGCATGCCGACCTCATCGAGCTCACCCGTCAACC	1388		1395 GIDALEN III SIII SIII SIII SIII SIII SIII SII	1415 ValAlaThrValLysGluAlaGlyArgSerIleHisGl	::: 7495 GGATCCCTTGCCCGTCCAGCCGACCGGCAATGTCATCG	1435 HisThrProGluLeuProLeuAlaProArg	GCACCAĊĊĠACCCGGGA	1445	1459 TYZABPThrGlyAlaSerThrThrGly		31ySe		1494 AlaABPALAAFGALABUULUALAYALUKYSIYYEETEETEETEETEETEETEETEETEETEETEETEETEE		7162 CGACTTGCAACGCGACAAGGCCCGGCTGCGCCCACAAG	1527 ProValIleValProGluLeuGlyLysProArgGlnSe	7117CCAGAAGTTGCGTATCCGAACCGGAC	1547 GlyAlaProPheAlaGlyHisLeuProArgGlySerPr	1567 ProArgLeuGlnGluGlySerLeuSerSerLysAl
OyPropro-ProproGlnAsnLeuGlnProGluSer-AspAlaProGlnGln 1014	8865	1015 ProGlySerSerProArgGlyLysSerArgSerProAlaProProAlaAspLysGluAla 1034	1052	8816 GCACCAGCGCGTTCCAACGCGCACGAACACACCAGGAACGTACCGCGATCTCGTCC 8757 OV	1057	8756 TCGCCAAGACCTGCCAACACCGACGACGTTCTCGAGCGTCCTCATCGAGCAAGCA		1086		GlyLeuHisAspThrAlaArgProValLeuProArgProProThrIleSer 1103	1123		Oy AlalleSerGlnGlyMetSerValGlnLeuHisValProTyrSerGluHisAlaLysAla 1143 Db	8474	1144 ProValGlyProValThrMetGlyLeuProLeuProMetAspProLysLysLeuAlaPro 1163 Db		1164 PheserGlyValLysGlnGlnGlnGlnGlagfirerProArgGlyGlyGlagaryFloriosiuser 1103 Db	1202		2	TCCGGCACGACGACCATGTCCCCGACAACGAGGGTCTGGCAAATCTGCC 8235 Oy		0y 1238 LeuTyrLysGlyThrIleThrArglleIleGlyGluAspSerProSerArgLeuAspArg 1257 Db	8137		CACGTCGTGATCGCCGGTAATGTGCAGAGCGGC	1277 1LeuSerTyrGluGlyGlyMetSerValThrGinCysSerLySGluAsportyArgseise 1297		
	DD 8924 GCCCC	Qy 1015 ProGly		Db 8816 GCACC			Oy 1058 DD 8696 GGCAC			1087	858 / 1104	Dh 8527 GCAGT	1124	Db 8482				DD 8413 ACGAG Ov 1184 LeuGl				Oy 1223			Oy 1258 GlyAr			Db 8094 GCIGA Ov 1297 rSer(

AGGTCTGGT----- 7118 SerProLeuThrTyrGluAspHis 1546 ACAACACATCTCGTAGCCGGGCCA 7070 ProvalThrMetArgGluProThr 1566 CTTCGTCAAACGCGTCGGCGAACA 7013 AlaSerGln-----AspArgLys 1584 GAGCATGGCACCGCCAGTTG--- 7319 ----SerLysLysHipAspVal 1473 rovalHisProLeuAspValMet 1493 CAGCTCAC-----CGA 7223 31uSer----- 1507 ::: ACGAACCCAACAACTCCCAGAGTC 7163 31yGlySerIleAlaArgGlyAla 1526 ---ProProSerArgAspLeuThr 1394 --AAACCCGCAACCACCTGGACC 7496 CACACGATCACCGCAGTCGGCCA 7376 :::::: ||| caccagccgagccgcarcccgca 7265 credececaccearacreedece 7658 CGTCGACACGATCGGCAGCCCTG 7598 eulysproAlaHisGluGlyLeu 1414 lulleProArgGluGluLeuArg 1434 GAGCACATCCCGATCACCGGAGA 7436 leThrGlnGlyThrProLeuLys 1458 rgSerTyrValGluAlaGinGlu 1368 ||||||:: | ACCGTGAGCCGCAGCAA 7712 lugiyThrProProPro--- 1387 |||||::: CGATTGCTGCGCATGAGCGGAAC 7831 1387 SerIleGluGlyLeuMetGlyArg----- 1333 ||::: ||SCATACGTGGGCAGCTCGACCTG 7951 ACCTCAGCACCACGCACATGGAT 7891 ...----GluArgHis 1340

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       This invention describes a novel method nucleic acid (I) and its encoded polypeptide (II) containing at least one region that encodes an enzymatic activity involved in biosynthesis of spinosyns. (I) are used (i) to identify, inactivate or modulate genes involved in the biosynthesis of cidentify, inactivate a library of polyketide synthases; (iii) for adding forosamine or trimethylthamnose to a spinosyn or polyketide asjnosms, which are used for production of the corresponding enzymes, which are used for production of the corresponding enzymes, uncluding production of transgenic plants that express (II) are markers for sequencing of the Sacharopolyspora spinosa genome. (II) are markers for sequencing of the Sacharopolyspora spinosa genome. (II) are markers for sequencing of the Sacharopolyspora spinosa genome. (II) are markers for sequencing of the sacharopolyspora spinosa genome. (II) are sense to raise specific antibodies, useful for identifying cypression clones in a gene bank. Cells transformed with (I) may produce (II) at significantly increased levels or produce new derivatives of (II). This sequence, ORF 21, encodes an S. spinosa polyketide synthase
                                                        2478
                             1072
                                                                                                                                    ----CATGTTT 1021
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives.
 pCysAsnArgArgThrProLeuThrAsnArgValTrp-GluAspArgProSerSerAlaG
                   lyserThrProPheProTyrAsnProLeulleMetArgLeuGlnAlaGlyValMetAlaS
                                                                                   ---GCAC
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                                                                                                                                       1065 GGGATGAACCGCCAGCGCGTGACCGAGGCCATGCTCCC---
                                                                                                                                                                                                                                                                                                                               S. spinosa DNA fragment encoding ORF21, SEQ ID 47
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P-PSDB; AAB70968.
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11149 CGGTGAGGCCGAACGCCCGGCAGACCAGTCCACCTGTGGAGTGGGCTTGTCCACGT
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                                                                                                                                                                                                                                                                                                                                                                                                             45 HisSerArgAspTyrAlaSerHisLeuSerProGlySerIleIleGlnProGlnArgArg
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ------SerGluAspLeuThrLysAsp---ArgSerLeuThrGlyLysLeuGlu
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      ProValSerProProSerProProHisThrAspProGluLeuGluLeuValProArg
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ArgProSerLeuLeuSerGluPheGlnProGlyAsnGluArgSerGlnGluLeuHisLeu
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Matches:
Conservative:
Mismatches:
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                                                                                                                                                         [ndel8:
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ସ୍ପ	3128 GCTTCGCCCGCGTGGAGGCCTCCTTCGCCCGGCCTGCACCCCGGGGCCG 3081	Ov 2197 SerProHisSerGluGlvG	SerProHisSerGluGlvGlvLvsArgSerProGluProAsr
රු සි	1872 GlnGlnargProSerValLeuHisAsnThrGlyMetLysGlyIleIleThralaValGlu 1891 ::: 3080 AGCCGCTGCGCCTCTGCCCAGACAACGTGCTAAAACGTGAGCACCCAGGCCCAGGC 3021	2135	GCTGCGCCGCGCGCGCGCCGCCGTGGTGGAAGCGGC
ð i	ProSerLysProThrValLeuArgSerThrSerThrSerSerProValArgPro	Qy 2217 GlyGlyGluAspGlyIleG Db 2090 GGCGCCGCGCGCGA	GlyGlyGluAspGly1leGluProValSerProProGluGly GGCGCCGCGGGGAGGG
8 & 1		Qy 2237 SerArgSerAlaValTyrProLeuLeuTyrArgAspGlyGlu	roLeuLeuTyrArgAspGlyGlu
8 &		Qy 2257 MetGlySerLye :: Db 2000 ATCGGGTGAGAAAGAGTTTATTTTCAGAGTGAGAAAATAAP	Serlye
8 &	2900 ACGCGGCCCCCCTTCGGGCACCGCGCGCCATGGGGGCTCGGCGCCCCGCTGC 2841 1932 uMetGluProValLeuLeuProLysGluAlaProArgValAlaArgProGluArgProAr 1952	Oy 2267 GlnPro-ProAlaPhePheSerLysLeuThrGl	AlaPhePheSerLysLeuThrG sAGATTTTTGGGGAGAGGGGG
음 샹	2840 GGCCCGTCTTCGTCTCTCTCTCGGCGGCCGGGGCCTCGCCGAGCTCCGGGGCCCG 2787 1952 gAlaAspThrGlyHisAlaPheLeuAlaLysProProAlaArgSe 1967		LysSerLysLysGlnGluIleAsnLysLysLeuAsnThrHi
ପ୍ର		DD 1880 CICIGGGGIGGIGGICGICITGIGGGGGGICCCIGG Qy 2303 uTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetPr	CICIGGGGICGGGGGGGGGGGGGGGGGGGGGGGGGGGG
දි දි	1967 rGlyLeuGluProAlaSerSerProSerLysGlySerGluProArgProLeuValProPr 1987	1823	 ggrgacggrccrccrccrgg
ð		Qy 2322ThrGly Db 1763 TGGGAAGAAGTGGATGGGG	ThrGlyLeuMetThrTyrArgSerGlnAl ::::: TGGGAAGAAGTGGGGGTCCTCCTCCTGGTGGGTCTTGG
8 &	<pre>2667 GTGCGCTGGGACTTTGCGCCCTTCGAGACCAGCGTCCGCGCCGCCGGCCGGC 2617 2007 aSerProAspProProAlaProProAlaSerAlaSerAspProHisArgGluLysThrGl 2027 </pre>	Qy 2337 rThrAsnMetGlyLeuGlu :: Db 1703 rCAGCAGATGGGGGTC	rThrasnMetGlyLeuGlualallelleArgLysalaLeuMe
qa (dececceratance	2353	
රු අ	2027 MSELY8ProPheSerIleGinGluLeuGluLeuArgSerLeuGlyYYrHisGlySerSe 2047	1649	CTTGGGGATGTTGGGGGTCCTTGATGATGATGGTGGTGGTGGTG
ð :	3luProValSerProValSerSerProSerLeuThrHisAs	Oy 2353	GGTCCCGGCGGGTCCTGGTGGATGTAGATGGTCCCGAGGG
8 8	2518 CGGCCGGCGCGCGGCCCGCTCGGGCCCATCAAGGTGGAGGCCATCTCGGACGA 2465 2067 pLysGlyLeuProLysHisLeuGluGluLeuAspLysSerHisLeuGluGlyGluLeuAr 2087	2354	TyrAspG] :::
qq	2464 CGA-GGAGGCCGAGGCCGGCAACCCCTGCTGCTGCTGCTGGGGGGGC 2412	Db 1529 GAGTTAGACGATGGTGGTC Qy 2362 0	GAGTTAGACGATGGTCGTCGTCATGGACTGCCCCATCT-GC O
දි දි	2087 gProLysGlnBroGly	1470	 GAGGCGCAGACGCTGCCGTGCATGCACAAGTTCTGTCTGGAC
Š		Oy 2370 oLeuAsnAla 	OLeuAsnAlaSerAlaSerLeuProAlaAlaMe
යි දි	2351 CCTCGAGGCCCTTCTCCTCCGTCTTCTCTCCCCTCGGCCCGGTCCTCGTCCTCG 2292 2117 rproleuleuGlnThrAlaproGlyVallysGlyHisGlnArg-ValValThrLeuAlaG 2137	2387 pGl	CleuThr
qa		Db 1364TCCTCCACCACGTGGA Qy 2396SerProGlyGly-	TCCTCCACCACGAGGACGGCTTCGTTTGTC ṢeṛProGlyGly
è 6	2137 lnHislleSerGluVallleThrGlnAspTyrThrArgHisHisProGlnGlnLeuSerA 2157 2232	1311	GGATGAGCCGGT
l & 1	laProLeuProAlaProLeuTyrSerPheProGlyAlaSerCysProValLeuAspLeuA	Oy 2407 Y	YArgProSerSerArgLyaAlaLy
8 &		Qy 2421 uAlaSerGlyAspArgPrc	uAlaSerGlyAspArgProProSerValSerSerValHisSe
qq	2169		

SeralaAsnalaPheAsnPr 2370 GluSerAsnSerAlaMetVa 2283 ::: | | | ::: | | GCGGCCATCTTGGCGGTGGT 1881 GlyLysAlaLysValSerGl 2407 LysSerProAlaProGlyLe 2421 HisAsnArgAsnGluProGl 2303 MetglyLyg----- 2353 GGGAGGTGGACGGTGTTGGT 1590 GTCCCGGTGGTCCCGGGTGG 1530 GlnTrpGluGluSerProPr 2362 GCCTGGACGTCGCGGCCACC 1471 TGGAGACGCCCGTGGAGGGG 1312 SerGluGly------As 2438 TTTATGGGCAGGTCTCGACG 1132 snLysThrSerValLeuGly 2216 lyMetThrGluProGlyHis 2236 ||::: GGCTCGGCGGCCGAGGATCG 2055 luGlnThrGluProSerArg 2256 ysSerProGlyAsnThrser 2266 ||| ::: GGTGGGAGGAAGAAGAGAG 1824 ProAlalleThrGly---- 2321 GAGGGTCTTGGTGGTGA 1764 AlavalGlnGluHisAlaSe 2337 GGTGGTGGTGGTGGTC 1704 GGTGGGAAGTAGAGGGT 1650 ----- 2353 MetprolleThrAlaAlaAs 2387 CCTCCA----- 1365 GC------GGC 2091 cccrrcrccrcrrcrcc 2001 AAGTTTGTGCTGTATTTTCT 1941

 	Qy 1560 alThrMetArgGluProThrP :::	1580	OY 1600 alprogluHisHisProHisP	Db 3872 TGCCCGAGCACCCGCGCGCGCGCGCGCGCGCGCGCGCG	Db 3822	Oy 1640 lylleProLeuAspAlaAlaA 	Qy 1656 laProAsnProThrTyrProF :::	1676		Qy 1696 isHisAsnThrAlaThrAlaN Db 3666GGGGCCGCGGCGCCCCG	Qy 1716 rgGluSerSerLeuAlaLeuA	Db 3608 CCGAGGAC		Oy 1756 sp-ArgLeuAlaTyrLeu	Db 3515 ACCCGCGCGCGGCGGCCTC: Ov 1768 PheSerSerArgHisSerSe	3464	Qy 1785 Leu	1790	3347	Db 3308 GCCGCCTCATCGTGCTGGAC	Qy 1818 ThrThrValGluHisAl	Db 3248 ACTACCACGICTACGICGGC		Qy 1852 AlaSerHisSerHisAlaHi	
luaspGlyArgSerSerGlyProProHisGluThrAlaAlaProLysArgT 		4992 AGCGTGACCCGCCGCTGCCGCACATCGGGGACGCCTTGGCCCCAGGACCCGCTCTGG 4933 1323 erSeralaSerIleGluGlyLeuMetGlyArgalaIleProProGluArgHisSerProH 1343	GCCCTGC	1343 isHisLeuLysGluGlnHisHisIleArgGlySerIleThrGlnGlyIlePro 1360	1 1	GCCGCGCGGACCCCCGCGCCGGGGAGGCGACCGTCGAGGCCCTTGGGCCCGGGGGCGCGCCCCCCGCGCCCCCCGCCCCCC	4755 GCCTTCGCCGCGCGCGCGCGCGCGCGCGCGGACCGGCGGCGGCGG	4695 GCCTGCGGGGGGTCCTCGAGGCGCTGCCGCGCTCCGGCTGCCGGCGCCCGCC	1397 TyrLygThrGlnAlaLeuGlyPro	1407 Leuly8ProhlaHisGluGly1413		4515 GCCCTCGCCGCACCGTGGCGCCGCTGGTGCGCTACAGCGTGGACGGGGCCCGCGCCC 4456	LeuvalalaThrValLysGluAlaGlyArg	4455 CGCGAGGCCGCCTGGACCTACGCCGCGCCCTCTTCGCCCCGGCCAACGTGGCCCCGGGCC 43%6	CGGCGGGGCCGGGGGCGGAAGCCCGCGGCCGGGC	1439 Leupro	146	4284 GCGGGCGCCCCTCCGGCCTCCGGGCCCTCCAGCCCCGCCAGCACCAAGTCC 4225	1462 GlyalaSerThrThrGlySerLysLysHisAspValArgSerLeulleGlySerProGly 1481	1482 ArgThrPheProValHisProLeuAspValMetAlaAspAlaArgAlaLeuGlu-Ar 1501		1501 gAlacysTyrGluGluSerLeuLysSerArgProGlyThrAlaSerSerSerGidyl 1520 1501 gAlacysTyrGluGluSerLeuLysSerArgProGlyThrAlaSerSerSerGidyl 1520 1501 gAlacysTyrGluGluSerLeuLysSerArg		4067 ACGAGGGGTCTCCGGGCTCTCCGCGGGGACGGCCCACGGCCACC 4020	1540 oLeuThrTyrGluAsphisGlyAlarrorneAlaciyhisLeuriOAlyGly-SelriOv 1900
λō	8 &	g &	g 63	රු සි	۲۵ ·	a ò	ብ &	qq	કે ક	8 8	qa .	දු පු	ò	셤 (£ 6	8 8	8 8	qq	λ _O q	à	QQ	상 원	ò	යු ,	Š

iisGlnHisSerProlleSerProArgThrGlnAspAlaLeu 1871 OHISLEUTYTPTOPTOTYTLEUIleArgGlyTYrProAspT 1676 CTCGGCGCTGCTCGCCCTTCGCCCACCGCC---- 3465 erSerPro-----LeuSerProGlyGlyProThrHis 1784 ----ThrLysProThr 1789 GGTGCTGCTGCTCTCGGCGCGGGACCTCGGCTTCGCCGGCG 3348 rgGluArgAspArgAspArgGluArgAspArgAspArgGlu 1809 ------cdcgc 3309 hr-----Ser 1817 || caccatcgaggactggcccgcggacggcccggccgtggggg 3249 ly-----GlyGlyGlySerSerSerArgPro 1851 HislleProLeuAlaPheAspProThrSerIleProArgG 1640 AhlaalaTyrTyrLeuPro------ArgHisLeuA 1656 creececrial recedences and second control 3750 gGlnThrIleIleAsnAspTyrileThrSerGlnGlnMetH 1696 aMetAlaGlnArgAlaAspMetLeuArgGlyLeuSerProA 1716 GCTCCGCCGCCGCCTGGATGCGGCAGATAGCCGACC 3609 JASnTyrAlaAlaGlyProArgGlyIleIleAspLeuSerG 1736 11.euValProProThrProGlyThrProAlaThrAlaMetA 1756 cracaccanaccanadacacccacaccaaccrada 3516 GCACGCCTGGGCCGGGAACTGGACCGGGCGCCCCGGACATCG 3408 ProArgLeuGlnGluGlySerLeuSerSerLysAlas 1580 SerThrProArgGluIleAlaLysSerProHisSerThrV 1600 -----crcegeeccececa 3873 ProlleSerProTyrGluHisLeuLeuArgGlyValSerG 1620 |||| |----- 3823 ...-----gagaccacacaccicade 3805 CCCGCGCTCGTCCTCCTCCTCCTCCTCCTCCG 3906 desecresesedes

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2455
                                                                              rSerAlaGlySerThrProPheProTyrAsnProLeuIleMetArgLeuGlnAlaGlyVa 2475
                                                                                                                                       7481 AGCACGATCACCAGCAACCCCACGGGCTAGCCTTCCCGAAATCGGGTCCGGTGTTACCG 7422
                                                                                                                                                                                                                                                                                                     Pseudorabies virus; PRV; LLV; large latency transcript; attenuated virus; vaccine; early protein 0; BPO; HSV-1 ICPO; protecting animals; deletion mutants; swine; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        New pseudorabies virus mutants for use in vaccine - having a deletion and/or insertion in the early protein O gene or large latency transcript
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                ORF2"
                                                                                           --GluGlyAspCysAsnAr
                  --ValTrpGluAspArgProSe
                                                                                                                      2475 lMetAlaSerPro---ProProProGlyLeuProAla--GlySerGlyPro---LeuAla
                                                          rcescreccioerceaecrceaegrearceaegreaaceaecriesaa.---criteric
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /*tag= c
/note= "encodes predicted amino acid sequence of
7014. .8425
                                                                                                                                                                                                                                                                                     DNA encoding Pseudorabies virus large latency transcript.
                                                                                                                                                                                                                                                                                                                                                                                                 from PRV strain InFh"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     /*tag= b
/note= "derived from PRV strain Ka"
8382. 8387
/*tag= f
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/note= "RNA cap site"
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1. 7013
/*tag= a
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                                         gArgThrProLeuThrAsnArg-
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                                                                                                                                                               GlyProHisHisAla 2497
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  2430 lSerSerValHisSer---
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                                                                                                                                                                                                                                                                                                                                                  virus.
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AAQ73500 shows the Pseudorabies virus (PRV) large latency transcript
(LLT). The basic sequence is derived from PRV strain InFh and FRV strain
(LLT). The basic sequence is derived from PRV strain InFh and PRV strain
(LLT). The LLT coverlaps and is transcribbed in the opposite orientation with
respect to the BPO (early polypeptide 0) and the immediately early gene
(IE180). EPO is nonessential for replicatio, LLT is the only gene
expressed during PRV latency, and the IE180 gene is absolutely necessary
for PRV replication. However there are 2 copies of IE180 in the genome.
It is expected that PRV lacking one of the IE180 copies is viable.
Deletions in the non-overlapping regions of these 3 genes will generate double deletion mutants. The invention is concerned with the
construction of attenuated viruses which have a reduced ability to
creativate from latency. This can be achieved by functionally disabling
the expression of the BPO gene, or by disrupting the synthesis of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                7208
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               7427
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     552 AspThrSerGlyGluAspAspAspGluLysGluAlaValAlaSer---LysGlyArgLys 570
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          571 ThralaasnSerGlnGlyargargLysGlyargIleThrargSerMetAlaasnGluala
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        630 HisGlyArgAsnTrpSerAlaIleAlaArgMetValGlySerLysThrValSerGlnCys
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                                                                                                                                                                                                                                                                                                                          Sequence 8438 BP; 1141 A; 2916 C; 3327 G; 1054 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            710 ValserGlyAsnGluGluGluMetValGluGluAlaGluAlaLeuHis-
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214
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811
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              Disclosure; Col 15-30; 43pp; English.
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	1271 luGlyLysLysGlyHisVal-LeuSerTyrGluGlyGlyMetSerVal 1286	11418 CAGCCGAACAGCGCAACATAGGCGTCCTGCGCCGGATTCACCTGCCGTAGCTGCGATGCC 11359	. 1287 ThrGlnCysSerLysGluAspGlyArgSerSerSerGlyPro 1300	11358 TCGCACTGTAGTCACGCAGCGCGGTAGAAATCGACGTGGCGAAACGCCGGCCAGTGCG 11299		11298 CCTCGGTGAACCACATCTCCGAGTAGGCGCTTTGCCACGAAGAACCCGGACAAGCGTT 11239			Pro	CCGAGATACCCTCGACGGCTCGACGAGTTCCTCCGCGGTGGCCCCGTTGGCGA 1111	1350 HislieArgGlySerliefnrGinGlyleProArgSerlyrvalGluAaGluGluAab 1209	GlvThr		ProPr		1401 AlaLeuGlyproLeuLysLeuLysProAlaHisGluGlyLeuValAlaThrValLysGlu 1420	11001 GCCGCGGGCCG 10990	1421 AlaGlyArgSerIleHisGluIleProArgGluGluLeuArgHisThrProGluLeuPro 1440	crccccdarcaaccca		10965CCCCGACGTCCGCACACTCCAGTGG-TTGGCCGGTGCGCAGATCTCTTCCACG 10913	146	10912 ACATCGGTGATGATCTCGATGAGTGCTGCAAGCTCGTCGGGATCGCGCTGCAGGTTTTCG 10853	AspvalArg	10852 GTGGACAGCAGATAGACGGTGGCCAGTTCGATGCCGGCTTCGTGGCACCACCGC 10799	SerLeulleGlySerProGlyArgThrPheProProValHisProLeuAspValMetAla	AGCATTTC-GGCGATCTTGGCCGCACCCATCCGGTAGCCGTAGCTGACGTCGTAGCTAGCTAGCTGACGTAGCTAGC	1495	10742 GCCGGCGCTGCGCCCATCGCCGGTTGCCGTCGCACACGCTATGTGCCCGGGGCAG	1515 AlaSerSerSerSerTivSerIleAlaArgGJyAlaProvallievalPro	10082 GICGGALITICGAGGCGGCCGAGGCGGCGGCGGCGGCGGCGGCGGGGGGG	1533 LeuGlyLyBProArgGinSerProLeuInriytsinAsphibsiyAlario-FineAl 	ואפו	10562	1
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1571	luGlySerLeuSerSerEstLysAlaSerGlnAspArgLysLeuThrSerThrProArgG 1591 :: AAGACGAGGCAACAGGCGAAATGAGCGGCCAG
1591	lullehlalupsSerProHisSerThrValProGluHisHisProHisProIleSerProT 1611 ::
1611	yrgluHisLeuLeuArgGlyValSerGlyValAspLeuTyrArgSerHisI 1628 ::: :::
1628	leproLeualapheAspProThrSerIleProArgGlyIleProLeuAspAlaAlaAla- 1647 ::: :::: Tccacgrotarrccgccggcacdgrogrogrogcrogrogrogrogrogrogrogrogrogrogrogrogrogr
1648	AlaTyrTyrLeuPro
1662	ProHisLeuTyrProProTyrLeuIleArgGlyTyrPro-AspThrAlaAl 1678
1678	aleuGluAsnArgGlnThrIleIleAsnAspTyrIleThrSerGlnGlnWetHisHi 1697
1697	
1717	uSerSerLeuAlaLe ::: GGCG
1734	
10061	GTGCGATCGCCGGAATCTTGCTGAAGATGTGTTGGCCGGCG
10001	Prohibbeutroval
1747	-ThrProGlyThrProAlaThrAlaMetAspArgLeuAlaT
1766	
1784	sLeuThrLysProThrThrThrSerSerSerGluArgGl ::: ::: ACGACGTTCGGCTACCACGAGTTCTTCCACGCCTGCACC
1804	
81	
1825	rolleTrpArgProGlyThrGluGlnSerSerGlySerSerGlySerSerGly 1842
1843	3S 1849
· 4*	erArgProAlaSerHisSerHisAlaHisGlnHisSerProIleSe

12515 GCCCGGCGCACCCCCGGCGCCGAA	1003 lnAsnLeu	12455 CACGGTTGCCTAACCCGCCGGTGCCGC	12395 CGGCGTCGCCGTACAGCCACCCGCCGG	1032 ysGluAlaPheAlaAlaGluAlaGlnL	12335 CGACACCGCCGGCACCGCCGGCACCGA	lyLeuPro	12276CCGGCACCACCACCGCCC	1072 roserAlaPheserTyrAlaProProG	1089	12167 CGCCGAGGCCGCCGGCACCGCCGGCGC	1094 roValLeuproArgProP	12107 cerrececcecceccidescec	1111 erAlaLysHisProSerV	1204/	1128 IYMECSETVALGINLEUHIBVALFIOI 1::::[]::: 11988 CCGTACGTACAG	1148 alThrMetGlyLeuPro	11958 CCGTCTTGTCCCGGCCCGCCGTTGGCG	1159 ysLysLeuAlaProPheSerGlyValL	::: 11898 TCGAAGCGAGGTTGATCGCGGCCAAC	1169GlnGluGlnLeuSerP	11838 TTGACCGCCTCCGCAGCCGCATAAGC	1185 lyValPro	11778 GCGTGAACGCCGACATTTGGGCGCTC	1191 luAlaSerValLeuArgGlyThrAlaL ::	11/18 AGGGAGGGGATTGCCGGGGAAACCTCA	120/ erileinruyaGilvilerioserinra	11658 GCCAACGCCGCGCGTTGGCCTGCGCC	11598 GCGCGCAACACGCGGGGCGCCACG	1.234 rollabenValleuTvriveGlvfhrT	11538 CAGCATCGGATGTCCCGCATGGATCAG	1254 rgLeuAspArgGlyArgG	11478 GGGTTTCACCTGGGCAGATTGCACAGC
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		13	735 GluCysSerGlyProAlaThrValAsnAsnSerSerAspThrGluSerIleProSerPro		13354 GGCGTTGCCGCCAACACCACCGCCCCATGGCGCCCTCACTGCCCGGAGGGGGT	755HisThrGluAlaAlaLysAspThrGlyGlnAsnGlyProLysProPro	13294 GGCGCCCTG	/ 771 AlaThrLeuGlyAlaAapGly-ProProProGlyProProThrProProArgArgThrSe 790	13234 ACCGTCGCCGAACAGCCCGGCGGTGCCGCCGCCACCACCGCCACCGCCGCACCGCCGCCGCCGC	790 rArgAlaProlleGluProThrProAlaSerGluAlaThrGlyAlaProThrProPro	61	131312 CGAGCCCAAGTTGCCGCCACGCCGCCGCCGCCGCCGCCGCCGCCGCGGA		13078 GCCGCCGGCACCACCATGCCGAACAACCACCCACCGTTGCC 13037		13	860 uProvalLysSerGluCysThrGluGluAlaGluGluGlyProAlaLysGlyLysAspAl :::	12977 TCCCGCCGACGCCGTTGCCGGGGTTGCCGAACAGCCCGCCGGCGGGCCCGGCCCGG	y 880 aGIUALAAIAGIUAIATARAIAGIUGIYAIALeuLysAlaGIULysLysGIUGIYGIYSE 900	900 rGlvaroalathrThrAlatvaSerSerGlvAlaProGlnAanSerAanSerAla-T	12875 GGCGAGGCCCCGAACAACAAGCGGCACGCCGCCGCCACGCACGCGCGA		12		12755 CGGCGCCGGCCGCCCTGACCGCCGGTACCACCGTCCCGCCGGATCCGGCGGCAC 12696	942	12695 CACCGCCGCCATACAGCCACCGGTGCCACCGGCGCCACCGGGGCCCCGGCCCCGG 12636		12635 CGGCGCCGCCGCCACCGGAAAGGCCACTTAAGCTACTACTGAAGGTGACGCTCTGAG 12576	968 lnArgAlaAla1leProProlleGlnValThrLysValHisG 983		783 INFIORIORIORIORIORIORIORIORIORIORIORIORIORI
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360GlyLeuSerMetSerAlaAlaArgSerGluHisGluValSerGluIleIleAspGl 378	378 yLeuSerGluGlnGluAsnLeuGluLysGlnMetArgGlnLeuAla 393 14203AAACTCCGCCAGTGGCCCTCGGTGCCGGAC 14172	394ValIleProProMetLeuTyrAspAlaAspGlnGlnArgIleLysPheIleAsnMe 412	tasnglvieuWetalaAspProMetLvsValTvrivsAspArgGlnValMetAsnMetTr			452 yLeuileAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyrLyrLe 472 14081 14081	472 uThrLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArgAr 490	g-Glybysserginginginginginginginginginginginging g-Glybysserginginginginginginginginginginginginging				542 LysGluAspLeuLeuLysGluLysThrAspAspThrSerGlyGluAspAsnAspGluLys 561 ::	562 GluAlaValAlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgArgLysGlyArg 581	IleThrargeerMetAlaAsnGluAlaAsnSerGluGluAlaIleThrProGlnGlnSer	13805 CTGACCGTTCG	cescecestracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearracearrac	622 ThralaLysGlyLeuLeuGluHisGlyArgAsnTrpSerAlaIleAlaArgMetVal 641 ::: 13718 CACCGGCACCGGTGCTGCGCTGGGCTGGCCGCCGT	642 GlySerLysThrValSerGlnCysLysAsnPheTyrPheAsnTyrLysLysLygLnAsn 661	LeuAspGlulleLeuGlnGlnHisLysLysRetGluLys			696 PheProProValValGluAspGluGluMetGluAlaSerGlyValSerGlyAsnGluGlu 715
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WP AA199682_43 4300001 4410000 WP AA199682_44 4400001 4411529	Alignment Scores: 1.09e-09 Length: 110000 Score: 618.00 Matches: 646 Percent Similarity: 30.90% Conservative: 267	Mismatches: Indels: Gaps:	US-09-522-753-5 (1-2517) x AAI99682_12 (1-110000)	Qy 150 ProProSerProProHisThrAspProGluLeuGluLeuValProProArgLeuSerLys 169	170		Db 14954 CTTGTGCCCCTATCGGCGATGCTCGCTGTTGTCGGAATTCGGGTTGGCCGCGCAGCC 14895 Qy 203 GlualaAlaLysProProGluBroGluLysProVal		Db 14834 GCGCCCCGCACCTGCCGCGCGATGCTCTGCAGCTGTGACGCATTGGCGGCCTCGGCGCCG 14775	Qy 215SerProProProlleGluSerLysHisArgSerLeuValGlnIleIleTyr 231		} .	Db 14687 GGCGTTTGCGTTGAACAGCCTAGCGATGGCGGCCAACACTTCATC 14643 Qy 267HisGluAsnIleLysIleAsnGlnAlaMet 276	CTAAC	<pre>Qy 277 ArglysLoulleLeuTyrPheLysArgAsnHisAlaArgLysGlnTip 294 </pre>	Qy 295	303AspGlnLeuMetGluAlaLeuGluLysLysValGluArgIleGluAsn ::: :::	GCACTTCAGACCGATGAATCACGTTC roArgArgArgAlaLysGluSerLys	14405 GACCCTATACGGTTTTALCACTGTACAACTAATTT	<pre>Qy 333 rGluLyeGluTheProGluTleArgLyeGluLeuGluGluGluGluGluGluGluGluGluGluGluGluGlu</pre>		Db 14309 AGCCGGGAAGCGTGGATCCCGTCGCTTCGATAGGCGGCGGGGAACGTAGGGGGATCGACGCG 14250

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                                                                                              "megalomicin 6-deoxyerythronolide B synthase 3"
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Qy 2426 GProProSerValSerSerValHisSerGludlyAspCygAsnArg 2441 Db 53321 TCCCGACCATGCCGGGGGCGCGGGGGGGGGGGGGGGGGG		Micromonospora megalomic Key compleme CDS compleme /partial /gene= " deoxyglu /note= " CDS 92820 /*tag= " /gene= " CDS /*tag= " /gene= " /product	CDS 33	FT /*tag= e FT /*tag= e FT /*tag= e FT /*gene= "megDII" FT / product= "TDP-3-keto-6-deoxyhexose 3-aminotransaminase" FT / note= "eryCI, DnrJ homologue, encodes AAB82205" FT / 58226595 FT /*tag= f FT //gene= "megDIII" FT //gene= "megDIII" FT //gene= "megDIII" FT //gene= "eryCVI homologue; encodes AAB82206"
54377 CATCACCCCGGCCCCGGCCCCGCGCCCCGCGCACCGCGCATCAGGAGGCTCGGC 54318 2134ThrheualaGlnHisIleSerGluValIle 2143 2134	ucllyMetThrGluProGlyHisSerArgSerAlaValTyrProLeuLeu :::	2778 SerAsnSerAlaMetValLyBSerLyBLyGGlnGluIleAsnLysLeuAsnThrHis 2297 :::		53501 AAGATCAGCCGCACGTACGTGCGTAGCCGGTCTTGAAGAAGTTCGAGTAGCCGGTGAGC 53442 2397 ProGlyGlyGlyGlybyBalaLy8ValSerGlyArgProSerSerArgLy8AlaLy8Ser 2416 23441 GCCTGCTCGTCGGCGGGGGGGGCCCCGCCCGAGCGCCTCGGCGCCTCCGGG 53382 2417 ProAla-Pro

		57077	Qy 1322 IleSerSerAlaSerIleGluGly, Db 57018	1335	56988	DD 56940 GGCGGCGTGC	Qy 1375 AlaLysLeuLeuLysArgGluGly	Db 56919 GGGCGCTTCAGCCCCGACCTGGC		Qy 1390 SerArgAspLeuThrGluAlaTyr	56799	Oy 1410 AraHisGludiyLeuValAlarini Db 56769 CCGGGTGGTGGAGGAGGACGACAG	Oy 1428 IleproArgGluGluLeuArgHis	Db 56709 GCGCCCGCTCTACGAGTTCGCCAC	Qy 1441 LeuAlaProArgProLeuLys-Gl	56649	Oy 1458 -LYSIYIASDIIICIYATASELIII	1472	Db 56529 CCACGAGACGGTGACCACCCTGC	Qy 1473 lArgSerLeuIle	Db 56469 CCGCACCGGGATCTGCGCCGACTC	Oy 1477	Db 56409 AGGCGAGCCACGCCGCAGAGCCGC 1478GlySerProG	56349 ACCACGCCGCAA	Qy 1493 tAlaAspAlaArgAlaLeuGluA:	Db s6295 GCTTGGCTCGTCCTGCTTGGCTC
992 hrlysproAlaProProAlaProProProProProGlnAsnLeuGlnProGluSer- 1009	1010AspalaProGlnGlnProGlySerSerProArgGlyLysSerArgSerP 1026	1026 ro		GTCCGCCCTCACCCCCGCCGGCTGGACGCCGTGGTGGCGCCCAAGGCGCT	1054 rophePro		57880 CTCCTCGGCGGCGGGGCTGCTCGGCACCCCGGCCAGGCCAACTACGGGGGGGG	1072 roseralaPheseriyrala	LeuGlyL		1088 euHibaspThralaargProValLeuProArgProProThrIleSerAsnBroProL 1108	eulleSerSerAlaLysHisProSerValLeuGluArgGlnIleGlyAlaIleSerGlnG	gc-cgcccccrgccgccgccacgccacgccacgccacgcc		TGCGCCAGCTCGTGCCCGTGCGGCGCAGGAGCGCCC	1143 AlaProvalGlyProvalThr	1154LeuprometAspprolysLysLysLeuAlaProPheSer 1165	57485 CGGAGCTGCTGCTGACGCTCGTGCGCCGCCACCGCCACCGTCCTCGGCCACCGGGGGC 57426	LeuGly		1186 ValProThralaGlnGluAlaSerValLeuArgGlyThrAlaLeuGlySerValProGLY 1205			1226 ArgGlySerlleThrHisGlyThrProAlaAspValLeuTyrLysGlyThr 1242		1243 IleThrArgileLieGiyoluAbbsetriosetArghedaspargicians
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TGAGCCGGGCGCGCGCGGGGCCGGGTCACGCAGAGG 56410 cedecertecreeercegerrrereceeecerrre 56236 ::::: GCTGGACCCGAGCACGCGCGCGTGAGCGCCCGAGAT 56470 GGGCCACGCTACGGAGGTGGACCACGCCGCAGAGGTGG 56350 ||||||| ------GCAGGTCCAGGGTGCGCTT 56770 AGGCACCTCGTGCTGCGCAGCCGCACCACCGACTG 56710 |||:::||| |GCCCTGGTGGGCAAGCGGCACCTCGTGGTCCGTGACGG 56530 scecaccerracecereracereceseses 56860 3ccggaccgrccractcgacccgarcaggcrgcccc 56800 acaccccagacgcrgrccaacrggacggcrrcac 56650 gThrTyrAspMetMetGluGlyArgValGlyArgAla 1321 -----GGGCGCATCTGCTTCGAGCTGAACCTGCT 56989 -----CGACGAGGA 56920 GlyArgThrPheProProValHisProLeuAspValMe 1493 ArgAlaCys---- 1503 -----GACGGCCCGTTCCC 57078 T------CGGCTTCACCACGACCGTGCTGTC 56941 3luGlySerIleThrGlnGlyThrProLeu----- 1457 G---GICACCCGCCCGACCGGGCCGCGCTGACCGCG 57150 rrysThrGlnAlaLeuGlyProLeuLysLeuLysPro 1409 ırValLysGluAlaGlyArgSerIleHisGlu----- 1427 ThrThrGlySerLysLysHis-------1471 lyThrPro---- 1384 -------ProProProPro------Pro 1389 rGluGlyLysLysGlyHisValLeuSerTyrGluGly 1282 rLysGluAspGlyArgSerSerSerGlyPro---Pro 1301 yLeuMetGlyArg-----Ala 1334 OHishisLeuLysGluGlnHisHisIleArgGlySer 1354 rTyrvalGluAlaGluAspTyrLeuArgArgGlu 1374

681 ArglyslyslyslysalaProAlaAlaAlaSerGluGluAlaAlaPheProProValVal 700

59217 desercecercaccececededeacecacecececerecerces

701 GluAspGluGluMetGluAlaSerGlyValSerGlyAsnGluGluGluMetValGluGlu 720

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rValSerSerValHisSerGluGlyAspCysAsnArgArgThrProLeuThrAsnArgVa 2449
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                                                                                                                                         CCCGACACCGCATCCCCACCCCCCAACACCCAAATCCCACAAACCCTCCGCACACACCACC 156
                                                                                                 lTrpGluAspArgProSerSerAlaGlySerThrProPheProTyrAsnProLeuIleMe
                                                                             tArgLeuGlnAlaGlyValMetAlaSerProProProProGlyLeuProAlaGlySerGl
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Novel maytansinoid produced by bacterial host cell transformed with expression vector comprising open reading frame from ansamitocin gene cluster I of Actinosynnema pretiosum, useful as antitumor agent.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       The invention relates to maytansinoid produced by bacterial host cell
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                                                                                                                                                                                                                                                                                     gene cluster I.
                   -------cárcgaacarccccgcgrcaracacaaacccc
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                                                                                                                                                                           155 CCACCCGGAAACGACAACCCATCCCCA 128
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                                                                                                            ThrValAsnAsnSerSerAspThrGluSerIleProSerProHisThrGluAlaAlaLys
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761 AspThrGlyGlnAsnGlyProLysProProAlaThrLeuGlyAlaAsp-----
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82746 545 171 877 743

Length: Matches: Conservative: Mismatches: Indels: Gaps:

7.34e-10 620.50 30.81% 23.45% 4.70%

> Percent Similarity: Best Local Similarity:

Query Match: DB: (1-82746)

JS-09-522-753-5 (1-2517) x AAL61224

		2162 oLeuTyrSerPheProGlyAlaSerCysProValLeuAspLeuArgArgProProSerAs 2182	AACGCCCGACACCCCCATCACCACCCCCCCCCGAGAAAACTCCACAAACACAACAACAACAACAACCACCACCAC	731 CCCGGCG		eAsnLysLysLeuAsnThrHisAsnArgAsnGluProGlulyrAsBillesErGliff		2369 nProLeuAsnAlaSerAlaSerLeuProAlaAlaMetProIleThrAlaAlaAspGlyAr 2389 ::: :::
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	2049 CGCATCCCGCACCACACCACCACCACATCCACCACCACCACCACCACCA			1929 yrProThrLeuMetGluProValLeuLeuProLysGluAlaFroArgValAlaArgProG 1949				1361 GTCCCCGACACCCCAAACGCCGACACCCCGC-ACGACGCACACCACCACCACCACCCAC

Š	1247 eGlyGluAspSerProSerArgLeuAspArgGlyArgGluAsp 1261	1521
q	3997 GCCCCGCCCCCCCCCAGGTGCCCGAAGAAAAACGAAGAAACGCTCCAAGGTCCAAGGTCCAAGGTCCAAGGAAAAACGAAAAAAAA	3091 GCAG
È	1261 1261	QY 1536 ProArgGinserFromeuinflyfGinabh
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ò	1262SerLeuProLysGlyHisVallleTyrGluGlyLysLy 1274	
qq	caccaacaaagrecrecascarasticascaaaaascaraacacaacaacaacaaag	1556
ර සි	1274 #GlyHisValLeuSerTyrGluGlyGlyMetSerValThrGlnCysSerLysGl 1292	Db 2917 CGGTGGTGTCGCCCCGTGGCGGGGTTGGT
ò	uAspGlyArgSerSerSerGlyProProHisGluThrAlaAlaProLysArg-ThrTyrA	1557
q	3758 CTCGGGTCGAGTCGATCAGGGCCACCCGGTCGGCCCGGCGGGGAGACCCG 3708	2857
ò	1312 spMetMetGluGlyArgValGlyArgAlaIleSerSerAlaSerIleGluGlyLeuMetG 1332	Qy 1565 ProfinrProArgLeuGinGluGLySerLeus Db 2797 CCAACGGGGGGCCGGCCCGGGGGGGGACACC
qq	ACGGTGGACGCGGCCGGGCC	1585
ඊ සි	1332 lyArgAlaileFroProGluArgHisSerProHisHisLeuLySGluGlnHisHisIlteA 1352 3684GGCGAGCACCACTGC 3669	Db 2737 ACGCCTCACCGACGGTCA
Š	rgGlySer1leThrGlnGly1leProA	
q	AGGCGGCCGGCAGGAGCGCGGCGACCGGGC	2695
ò	1361 rgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGluAlaLysLeuLeuLysArgG 1381	Oy 1624 rArgSerHislleProLeuAlaPhe II
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ò	1381 luGlyThrProProProProProSerArgAspLeuThrGluAlaT 1397	Db 2608 GATGCCCGGAACACTC
qq	3599 Ardeneccedecedecedecedecededededesegededegededededer 3540	1656
è 1	yrLysThrGlnAlaLeuGlyProLeuLysEvoNlaHisGluGlyLeuValAlaT	
a é	CGACCGTAGACGCCGGGGGCCCGTACGGCGGCGGCGGCCTGGTCCTCG	Oy 1676 rAlaAlaLeuGluAsnArgGlnThrlleIle
දි දි	141/ nEvalLygGildataGiyArgGsEllenisGillierroArgGstucilleurghiskiir 143/ 3497 CCCTTCGRAGCCGCCCACCATCGACGACGCACGAACGCACAACGAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACATCAACAA	Db 2530 CACGCAGATTCCGATACCAATACCCACCATC
8 8	LeuProLeuAlaProArgProLeuLysGluGlySerIleThrGlnG	1696
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ò	1454 lyThrProLeuLysTyrAspThrGlyAlaSerThrThrGlySerLysLysHisAspV 1473	QY 1/06 GALAASPMetLeuargGIyLeuserProArg Db 2410 ACAGCATCCCTTCCACGCTTCCACACATGCG
qq	ceaccegeccacacacacacacacacacacacacacacac	1725
8	al-ArgSerLeulleGlySerProGlyArgThrPheProProValHisProLeu	Db 2350 GCCGCACCACCCCATCCCGCTCACACTC
8 8	3331 CGGCGACGCCCCCCGGGTGACGCCCCCAGAGGCGGGCTCCCCCCGCGTCCGCGTCCGCGATG 3272 1491 AsovalMetAlaAsoAlaArGAlaLeuGluArGAlaCvsTvrGluGluSerLeuLvsSer 1510	GlyT
g 43	ccrgarcaggcgagcgagcggrcaggcgagcgagcggggcggcggcgcgcccg	2291
ò	1511 ArgProGlyThrAlaSerSer	OY 1764 aPrOGINPROPHESERSETATGHIBSETSET DD 2258 ACCCCCCACCCAACCCAACCGACCCCACCCCACCCCAC
qa .	cácicireaceadéceageagaceaceaceceaceageceaceaceaceaceaceaceaceaceaceaceaceacea	1784
රු දි	1518	Db 2198 GGCAGTCCGACCACCATCC
3		Oy 1804 gAspArgAspArgGluArgGluLysSerIle

1521	GCAGTICGAAGGCGCCACGCCAGCCAGCCTCCCTACGCCCCCCCACAAAABAAGGAAGCGCCACGCCA
1536	ProArgGlnSerProLeuThrTyrGluAspHisGlyAla
1549	GGTCGTCGGTCGCAACGGTTTCCAGGTGATCCGATACCGCAGGTCACCCGCCGGGTCG 2918
1556	1556
2917	CGGTGGTGGTGTCGCCCGTGGCGGGGTTGGTGGCGGGGGTTGGTGG
1557	GGTTGGTGGGAGTCGGTGCCCAGTAGTGGCGGCGTTGGAACGATACGTCGGCAGGTCAA 2798
1565	ProThrProArgLeuGlnGluGlySerLeuSerSerSerLysAlaSerGlnAspArgLys 1584
1585	LeuThrSerThrProArgGluIleAlaLysSerProHisSerThrValProGluHisHis 1604
09	162
62	63
1636	rlleProArgGlyIleProLeuAspAlaAlaAlaAlaAlaTyrTyrLeuProArgHisLeuAl 1656 :::
1656	aProAsnProThrTyrProHisLeuTyrProProTyrLeuIleArgGlyTyrProAspTh 1676
1676	ralaalaLeuGluAsnArgGlnThrIleIleAsnAspTyrIleThrSerGlnGlnMetHi 1696
1696	BHIS
1706	GAlaAspMetLeuArgGlyLeuSerProArgGluSerSerLeuAlaLeuAsnTyrAl 1725
1725	aAlaGlyProArgGlyIleIleAspLeuSerGlnValProHisLeuProValLeuVa 1744
74	176
1764	aProglnProPheSerSerBrgHisSerSerSerProLeuSerProGlyGlyProThrHi 1784
1784	SLeuThrLysProThrThrThrSerSerGluArgGluArgAspArgAspArgGluAr 1804
Ó	gAspArgAspArgGluArgGluLysSerIleLeuThrSerThrThrValGluHisAl 1824

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Novel isolated polypeptide involved in biosynthesis of macrolides by microorganisms, useful for biosynthesis of macrolides by microorganisms, preferably for biosynthesis of rosaramicin.
                                                                                                                                            AsnArgArgThrProLeuThrAsnArgValTrpGluAspArgProSerSerAlaGlySer
                                                                                                                                                                CCGGCGTCTGCG------CGCCCTGTCCGGTGAACAGCA
                   ----GlyGlyLysAlaLysValSerGlyArg---
                                                                                                    ---SerGlyAspArgProProSerValSerSerVal-----HisSerGluGlyAspCys
                                                                                                                         17913 cecaccédrogacectrocogaacecrescaacecrescacecacacacecces
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                                                            ------ProSerSerArgLysAlaLysSerProAlaProGlyLeuAla-----
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/note= "CDS does not include sti
/partial
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The invention relates to genes and proteins involved in the biosynthesis of macrolides by microorganisms. In particular it relates to the nucleic acids forming the biosynthetic locus for rosaramicin (a 16-member macrolide antibiotic) from Microemonospora carbonacea. The invention is useful for the biosynthesis of macrolides by microorganisms. It allows direct manipulation of macrolides and related chemical structures by chemical engineering of the proteins involved in the biosynthesis of rosaramicin. It is useful to catalyse certain biochemical reactions, in vitro or in vivo, to direct or enhance the synthesis or modification of a polyketide substrate or its precursor. The present sequence is M. carbonacea polyketide synthase (PKS) type I gene. (Updated on 27-OCT-2003 to standardise OS field)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        GlnHisHisSerArgAspTyrAlaSerHisLeuSerProGlySerIleIleGlnProGln 62
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Sequence 11238 BP; 1110 A; 3596 C; 4707 G; 1825 T; 0 U; 0 Other;
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1995 CATCCTTCCACCACCACCCCCCCCCCCCCCCCCCCCCC	18210 CGCCGCCCGCCGAGAGCCCTCGCATCAGCCGCGCGCGCCGCCGCGCGCG
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qq	21701 GCGGGCTTGCCCGTACACGTCGCTCAGCGCCCGCACCTCGATCGGGTCGCCCAGCGCCGT 21642	è	1472 AspVal
ò	1184 uGlyValPro	3 43	20760 GGTCCI
qq	21641 CCCGGTCCCGTGGCATTCCACGAAATCGACGTCGGCCGGAACCACCCCTGCGTGCG	õ	1487 ValHis
ò		qq	20703 CTGTCC
셤	21581 CGCCTGGCGTAACAGCGCCTCCTGGGCAGGGCCGCTGGCAGGCCCGCTGCTGGTCGG 21522	λ	1507 SerLev
ò		q	20682TTC
qa	21521 GCCATCATGATTGATCGCCGTACCCCGGATCACCGCCAGGATGGGGTCGCGGTCGCGCTG 21462	à	1527 ProVal
ò		qq	20631 CCACC
ą G	21461. CGCGTCGGAGAGCCGCTTGAGCACCÁCCACGGCGCAGCCTCGGCCCGCGGGGTAGCCGTC 21402	ò	1545 AspHis
ò		q	20571 GCTCAC
qq		È	1560 ValTh
ò		đ	20526
qq	21353 CATCCGTGAGAGCAGCGCGAAGGTCTTCGGCGAGCAGCAGCATGTTGACCCCGGCCAACCAG 21294	λō	1580 SerGl
λõ	g-LeuAspArgGlyArgGluAspSerLeuProLysGlyHisVallleTyrGluGlyLysL	qq	20487 TGCCT
đ	CTGGCAGCC	ò	1600 ValPr
ò		q	1 20430 GCGCC
qq	21254 GAGGTGCAGCGCCACGAGCGACGAGCAGCACGCGTATCCAC 21213	λõ	1620 GlyVa
à		g	20391 GCTCT
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ò	1312 spMetMetGluGlyArgValGlyArgAlaIleSerSerAlaSerIleGluGlyLeuMetG 1332	q	 20331 AGAAG
q _C	21152 CATGITGCCGGTGCCGCTGT	ò	1659 ProTh
ò		đ	20280 CCATC
qa		λõ	1679 LeuGl
à	1346 LysGluGlnHisHisIlehrgGlySerIleThrGln 1357	d d	20253 CGCGC
QQ	21074 GGGGCCCAGGAACACGCCCGTGGGGCTCTCGCGCAGGCCGACGGGTCCTGGCCCGCGCG 21015	ò	1699 ThrAl
ò	alG	q ₀	20208 TCGGC
qq	21014 CTCGATCGCCTCCCAGTCACTTCCAGCAGCCGCTGTTGCGGGTCCAGGCT 20961	λ0	1719 SerLe
ò		qa	20193
дg	20960 CATCGCCHC	č	1739 HisLe
'n		qq	:: 20181 CATGT
qq	20930 CGCCGCATCGAACGICTCCACCTCGCCACAAAGCCACCCTGGGCACGTAGGTCTGTCT	8	1759 AlaTy
ò	HisGluIleProArgGluGluLeuArgHis	q	20139
q _Q	20870 CGGAGCCTCTCCCGAGCCGGGGCCGCGCCCGTCTGCCCCAT 20830	ò	1779 ProG
ζ	ThrproGluLeuProLeuAlaProArgProLeuLysGluGlySerIle	q	20094 GACGC
qq	20829 TCCACCGGTCGGCGCACCTCGGTGCTGACCACCACGCCCT	ò	1794 SerG
δ	ThrGlnGlyThrProLeuLysTyrAspThrGlyAlaSerThrThrGlySerLysHis	dg	1 20034 TCGA(
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Series S	1472 AspVal 20760 GGTCC7 1487 ValHis 20703 CTGTCC 1507 SerLev 20682TTV 1557 Prova 1558 Prova 1550 ValTh 20526 1560 ValTh 20526 1560 ValTh 20430 GCGCC 1653 ArgGl 20431 AGAAC 1659 Prova 1779 Prova 1770 Prova	1486 2070 1506 2068 1526	TTCGCACGTCGGTGCGTCCAGCGCCTGGCTCAGCAAGTATT 20632 ProVallleValProGluLeuGlyLysProArgGlnSerProLeuThrTyrGlu 1544	ArgGlulleAlaLysSerF ::::: :CGGCGACCACGCGCGCACACACACACACACACACACACA	:::	ovalLeuvalpr cggcgcrgcgc cggcgcrgcgc
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Oy 257 rAsm Db 24384 GACG Oy 277 gLysi Db 24339 GGGG Oy 297 8Phe		Db 23605 ¢Ġ-
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epoE gene "encodes modules 7 and 8" "ancodes KS7" "4626 "encodes AT7" "41922 "encodes KR7"	rity: 5.73e-10 Hength: 660 rity: 31.32 Marches: 660 rity: 31.32 Marches: 1065 illarity: 22.35 Marches: 1065 illarity: 22.35 Marches: 1065 illarity: 22.35 Marches: 1065 illarity: 22.35 Marches: 1065 indels:	lyerogi
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257	rAsnGlnProSerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetAr 277
277	gLysLysLeuIleLeuTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLy 297
297	sPheCysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGluArglleG1 317
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337	eProGluIleArgLysGlnArgGluLeuGlnGluArgMetGlnSerArg 353
354	ValGlyGlnArgGlySerGlyLeuSerMetSerAlaAlaArgSe 368
368	rGlu
382	
402	aAspGlnGlnArgIleLysPheIleAsnMetAsnGlyLeuMetAlaAspProMetLysVa 422 :: CAGCCGTGCCGCGTG
422	ITyrLysAspArgGlnValMetAsnMetTrpSerGluGluGluLysGluThrPhe 440
441	
451	eGlyLeulleAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyrTy 471
471	rLeuThrLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArgArgGl 491 ::
491	. ytysserginginginginginginginginginginginginging
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529) ladluLysGluGluLysProGluValGluAsnAspLysGluAspLeuLeuLysGluL 549
549	
566	6 erLysGlyArgLysThrAlaAsnSerGlnGlyArgArgLysGlyArglleT 583

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encodes ketoreductase (KR) 2, the KR domain of
                                                         "encodes ACP2, the ACP domain of module 2" 37907
                                                                                       epoD_gene
"encodes modules 3-6"
.17606
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.22082
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.31373
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"encodes KR3"
.20642
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.30869
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.32807
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24647
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.25151
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27883
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.34067
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                                        *tag= f
/note= "encodes enoyl reductase (ER) of the loading
domain, potentially involved in formation of the thiazole
                                                                                         *tag= g
note= "encodes acyl carrier protein (ACP) of the loading
                                                                                                                                                                                                                                                                                                                                                                  *tag= q
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861. .8905
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090. .9179
                                                                                                                                                                                                                                                                                   *tag= n
note= "encodes adenylation domain Al of the NRPS module"
898. .7921
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note= "encodes adenylation domain Al of the NRPS module"
261. .8308
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966. .8983
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183. .9992
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note= "encodes condensation domain C7 (partial) of the
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898. .7921
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0121. .10138
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odule 2"
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note= "encodes thiolation domain (PCP) of the NRPS
odule"
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note= "encodes adenylation domain A10 of the NRPS
nodule"
.0261. .10306
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3327. .13899
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label= epoC gene
note= "encodes module 2"
0654. .12033
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label= epoB_gene
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Qy 2407 GlyArg	Db 19723 CGATCGGGGGTTTGTGGCCCATGGTTCACGACGCGCATGGCACCACGCTCCCCGAA 19664 Qy 2437 GlyaspCysAsnArgArg	UL 29	15-SEP-2003 (re- 12-SEP-2000 (fil- Sorangium cellul- Epothilone; poly- epol; epok; P450 tubulin polymeri Polyangium cellul	FT CDS 3.992 FT CDS 3.992 FT CDS 4.1291 A PT	FT /label= epoA gane /label= elabel= e
sserg		Db 20612 CAGACCGAGCCCACCAGACCGCCGGTCACAAGGTAGGTGCTGTCCGCGGGGAT 20559 Qy 2199HisSerGluGlyGlyLyBArgSerProGluProAsnLysThrSerValLeuG 2216		2309 roGlyThrGluIlePheAsnMetProAlaile	Oy 2365

원 수 명

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	-	848	qu
	N	24868	ò
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		24808	ò
	1052 yLeuproPheproValProProArgGluValIleLysAlaSerProHisAlaProAspPr 10	1072	QQ
	oSerAlaPheSerTyrAlaProProGlyHisProLeuProLeuGl	1087	કે દ
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		1142	3 8
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	1142 sAlaProvalGlyProvalThrMetGly	1155	qa
	oMetAspProLysLysLeuAlaProPheSerGlyValLysGlnGluGlnLeuSerPr	1174	දි දි
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	1174 oArgGlyGlnAlaGlyPro	1186	ਰ 연 전
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	GlySerIleThrLyBGlyIleProSerThrArgValProSerA	1220	q
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22955 regggargegeregreerelagegaaagegergeerreareageeeegergeerg 22896 23286 23285 İCGCCAĞĞCTGAACĞCCAĞTCCCCGAĞCCCĞAĞCTCCĞĞĞTĞCATĞTCCAĞĞTĞCTĞĞ 23226 23509 CTCCCGGAACGCTGGCCACGCGCGAAGCCCC-CGGCCATGCCCGGCGTCTGCGCGC 23451 23391 1522 IleAlaArgGlyAlaProVallleValProGluLeuGlyLysProArgGlnSerProLeu 1541 1542 ThrTyr-----GluAspHisGlyAlaProPheAlaGlyHisLeuProArgGlySerPro 1559 1560 ValThrMetArg-----GluProThrProArgLeuGlnGlySerLeuSerSerSe 1577 -----GlulleAla-----ThrValPr 1601 23731 GGCCACCAGGTCCCGATGCTATGCCCAACCAGGAGCTCCGGGTCTACGCCCCACGACCG 23672 1459 1502 AlaÇyşTyrGluGluSerLeuLysSerArgProGlyThrAlaSerSerSerGlyGlySer 1521 1426 -------HisGlulleProArgGluGluLeuArgHisThrProGluL 1439 1459 yrAspThrGlyAlaSerThrThr-GlySerLySLySHisAspValArgSerLeulleGly 1478 23791 roProGluArgHisSerProHisHisLeuLysGluGlnHisHisIleArgGlySerIleT 1356 1356 hrGlnGlyIleProArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGluAlaL 1376 1376 ysLeuLeuLysArgGluGlyThrProPro-----1385 1386 -----Throproproproperty 1397 1397 yrLysThrGlnAlaLeuGlyProLeuLysLeuLysProAlaHisGlu------ 1412 1316 lyArgValGlyArgAlaIleSerSerAlaSerIleGluGlyLeuMetGlyArgAlaIleP 1336 1296 erSerSerGlyProProHisGluThrAlaAlaProLysArgThrTyrAspMetMetGluG 1316 rLys-----AlaSerGlnAspArgLysLeuThrSerThrProArg------1439 euProLeuAlaProArgProLeuLysGluGlySerIleThrGlnGlyThrProLeuLysT 23330 ACGTCACCGCCACCGCGGAGCCGGTGGCTCATCGCGC------TGCGCGTCG -----PheproProValHisProLeuAspValMetAlaAspAlaArgAlaLeuGluArg 1413 -----GlyLeuValAlaThrValLysGluAlaGlyArgSerIle------23611 GTCGAGCAACAACGACTCGGCGTCCCCCCCTCCGCCCACATCA----23793 CG-------1577 1591 1479 1484 1336

26006 AGCGCTCCACAGCTGGCACGCCGAGGAGGCCTCTTTCGTGGCAGTTACCAGCCGAT 25947 769 ProProAlaThrLeuGlyAla			940
8 8 8 8 8 8 8	8686686	8686868	86868686868
	26761	638 AlaArgMetValGlySerLy8	676 GluargasnalaargargLysLysLyslalaaProalaalaalascrGluglualaala 676 GluargasnalaargargLysLysLysLyslalaaProalaalaalascrGluglualaala 671 GluargasnalaargargLysLysLysLyslalaalaalaalaacrGluglualaala 6710 GluargasnalaargargCGGTCCTGCAGCGCTGGCTCAGCAAGT 6710 GluglugluwetValQluglualagGlugluwetGlualascrGlyValSerGlyAsn 6714 GluglugluwetValGluglualagGlualateuHisAlaScrGlyAsnGluValPro 6715 GCAGCTCACCCTGAAGCGGTTGGGAATCTCCAGGGCGTTGGGGATGGTGGTGGTGGTGGGTTGGGGATGGTGGGGGTTGGGGATGGTGG
8 8 8 8 8 8 8	3 4 8 4 8 4 8 8 8 8 8 8 8 8 8 8 8 8 8 8	8 6 8 6 8 6 8 6	6 B 6 B 6 B 6 B 6 B

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cc proteins involved in the biosynthesis of epothilones. Epothilones A and B are 16-membered macrocyclic polyketides with an acytcysteine-derived starter unit; polyketides being synthesised from two-carbon building cc blocks, the beta-carbon of which always carries a keto group. Each round of two-carbon addition is carried out by a complex of enzymes known as the polyketide synthase in a manner similar to fatty acid blosynthesis. C EPOS AAYYSB573 and EPOS B, BPOS C, EPOS D and thiazole ring formation of epothilones, and EPOS B, EPOS C, EPOS D and C thiazole ring formation of epothilones, and EPOS B, EPOS C, EPOS D and C EPOS E (AAYSB579) is an epothilone macrolactone oxidase, and the proteins or an epothilone macrolactone oxidase, and the proteins or a (AAYSB575-YSB579) is an epothilone macrolactone oxidase, and the proteins or substituted for taxol in cancer chemotherapeutic compositions.

CC FORT (AAYSB575-YSB579) is an expelling a much lower drop in potency against a multiply drugtransport. Epothilones mint taxol, and are considerably less efficiently exported from such calls by the multidrug resistance protein (MDR, or P-glycoprotein). Despite the potential of epothilones are too complex for industrial scale chamical synthesis, and compliance allosem is difficult to ferment or producing poor yields of epothilones. The nucleic acids of the invention may be used for the ecombinant production of epothilones in a heterologous host that is more amenable to fermentation. (Updated on 15-SEP-2003 to standardise OS estandardise OS
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Sequence 68750 BP; 9596 A; 22458 C; 25537 G; 11159 T; 0 U; 0 Other;

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US-09-522	-753-	5 (1-2517)	x AAZ55887 (1	-68750)		
λΌ	13	ArgalaThr	GluProArgTyrProP	roHisSer-LeuSer	ArgalaThrGluProArgTyrProProHisSer-LeuSerTyrProValGlnIleAlaAr	32
qq	28316	CGAGCGACA	cadecadescarcacadecececedadaacecerdearcadecec-	cgagagccctdcar	cagccccgcgcc	28263
ò	32	gThrHigTh	qThrHisThrAspValGlyLeuLeuGluTyrGlnHis	GluTyrGlnHis	HisSerArgAspTyrAl	20
qq	28262	CCGCCACGA	ceccaceaeccrcaccccarcrrcaeceaeaacacccccccc	CAGCGAGAACACCC	cgccacgcacgc	28211
λ̈́o	20	aSerHisLe	uSerProGlySerlle	IleGlnProGlnArg	aSerHisLeuSerProGlySerIleIleGlnProGlnArgArgArgProSerLeuLeuSe	70
QQ	28210	-ceccaccaecreeced	GCTCCCCGGCGCTATG	GCGCTATGCCCAGCCACCAGCTCCGG	cggcrcra	28164
'n	70		nProGlyAsnGluArg	SerGlnGluLeuHis	rGlupheGlnproGlyAsnGluArgSerGlnGluLeuHisLeuArgProGluSerHisSe	06
οp	28163	CGCCCCACG		CAGCGCGTACTCCAC	CGTGAA	28118
δ	90	rTyrLeuPr	oglubeuglybysser	GluMetGluPheIle	rTyrLeuProGluLeuGlyLysSerGluMetGluPhelleGluSerLysArgProArgLe	110
qq	28117	1	gagcgcgggcrgggrgaacgccgrcrggrcgagcaacaacaa-	GAACGCCGTCTGGTC	GAGCAACAACGA	28076
λ̈́o	110	uGluLeuLe	uProAspProLeuLeu	ArgProSerProLe	uGluLeuLeuProAspProLeuLeuArgProSerProLeuLeuAalaThrGlyGlnProAl	130
qq	28075	CT	ceececrcccceecrc	CGCCCACATCACCTC	-cresececrecessercescentares established	28024
٥٨	130		luAspLeuThrLysAsL	ArgSerLeuThrGl)	aGlySerGluAspLeuThrLysAAspArgSerLeuThrGlyLysLeuGluProValSerPr	150
qq	28023			cccdd	cccddrcaacagcgcdacgcaccgg	27997
ò	150	oProSerProProHi	COPTOHISTATABDPTC	oGluLeuGluLeuVa	SThrAspProGluLeuGluLeuValProProArgLeuSerLysGl	170
g	27996		TCGAACGCCTCCCGGAACG	c r idac	-crescecececececana ecce	27952
ò	170		leGlnAsnMetAspArg	gValAspArgGluIl	uGluLeulleGlnAsnMetAspArgValAspArgGluIleThrMetValGluGlnGlnIl	190
Д	27951		CCCATGCCCGGCGTCTGCG	TOTOCCETOR.	-cgcccrgrccgcrgaacaggaacgccagc	27904

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						396 rôproMetLeuTyrAspAlaAsp	396 robroMetLeuTyrAspAlaAsp	396 ròproMetLeuTyrAspAlaAsp	356GlnArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluH 370		338 rodlulleArgLysGlnArgGluLeuGlnGluArgMetGlnSerArgValGly 355	
13.06 TGCCCGTACACGTCCCCCCGCATCGGTCGCCCCAGCGCCCTCCCGGTC 2724'	TGPTCGGTACACGTCGCACGCACCTCGATCGGGTCGCCCAGCGCCCGGCCGG	13.06 TGCCGGTACACGTCGCGCACCGCACCGGTCGCCCCAGCGCCGTCCCGGTC	136 TGCCGGTACGCGCTCAGCGCACCTCGATCGGTCGCCCAGCGCCCTCCCGGTC 2724 1	13.06 TGCCCGTACACGTCCAGCACCTCGATCGGTCCCCAGCGCCCTCCCGGTC 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724	13.06 TGCCCGTACACGTCCAGCACCTCGATCGGTCCCCAGCGCCCTCCCGGTC 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724 2724	336 TGCCGGTACACGTCCAGCACCTCGATCGGGTCGCCCAGCGCCTCCCGGTC 2724 336 TGCCGGTCCCAGCGCCTCCCGGTC 2724 336 TST.	1306 TGCCGGTACACGTCGCTCAGCGCACCTCGATCGGGTCGCCCAGCGCCGTCCCGGTC 2724	1306 TGCCGGTACACGTCGCTCAGGCACCTCGATCGGGTCGCCCAGGCGCTCCCGGTC 2724 396 TôProMetLeuTyrAspAlaAsp	356GlnArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluH 370		338 rodlulleArgLysGlnArgGluLeuGlnGluArgMetGlnSerArgValGly 355 7466	
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product= "Type I polyketide synthase, EPOS D (AAY58577)"
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*tag= k
                                                                    product= "Type I polyketide synthase, EPOS E (AAY58578)"
            "Type I polyketide synthase, BPOS C (AAYS8576)"
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/note= "No termination codon given in the specification"
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(AAX58579)"
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P-PSDB; AAXS8573, AAXS8574, AAXS8575, AAXS8576, AAXS8584, AAXS8585,
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AAXS8586, AAXS8587, AAXS8588, AAXS8590, AAXS8591, AAXS8592, AAXS8593,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               This sequence represents a 68.75 kb contig from Sorangium cellulosum comprising 22 open reading frames (ORFs) and includes genes encoding
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New isolated epothilone synthase genes, used for the recombinant production of epothilone for use in cancer therapy.
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product= "Orf 14 protein (AAY58593)"
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complement(65008. .65874)
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complement(64287. .64727)
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complement(63853, .64290)
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complement(65871. .66338)
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24-SEP-1998;
05-FEB-1999;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            23-DEC-1999
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                AAY58594.
                                                                                                                                                                             CDS
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 38346 CCCGCCGGTGCCGGCGTCGCTTGCCC------CGCCG-GCGCCACCGACCCGCTT 38395
                                                                                                                                        2436 luGlyAspCysAsnArgArgThrProLeuThrAsnArgValTrpGluAspArgProSerS 2456
                                                                                                                                                                                        2456 erAlaGlySerThrProPheProTyrAsnProLeuIleMetArgLeuGlnAlaGlyValM 2476
                                                                                                                                                                                                                                     2476 etAlaSerProProProProGlyLeuProAlaGlySerGlyProLeuAlaGlyProHisH 2496
2382 ProlleThrAlaAlaAspGlyArgSerAspHisThrLeuThrSerProGlyGlyGlyGly 2401
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       product= "Type I polyketide synthase, EPOS A (AAY58573)"
1872. .116104
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               product= "Type I polyketide synthase, EPOS B (AAYS8575)"
21746. .43519
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /product= "Partial Orf 1 protein (AAYS8S80)"
/note= "No initiation codon given in the specification"
complement(1900. .3171)
                                                                                                                                                                                                                                                                                                                                                                                                                                                               Epothilone biosynthesis; type I polyketide synthase; taxol substitute; anticancer; ds.
                                           2402 -------LysAlaLysValSerGlyArgProSerSerArgLysAlaLysSer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /*tag= g
product= "Non-ribosomal peptide synthetase, EPOS
(AAY58574)"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /product= "Orf 4 protein (AAYS8583)"
6226. .6675
/*tag= e
/product= "Orf 5 protein (AAYS8584)"
7610. .11875
/*tag= f
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                3415. .5556
/ tege c
/product = "Orf 3 protein (AAYS8582)"
complement (5612. .5992)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     product= "Orf 2 protein (AAY58581)"
                                                                                                                                                                                                                                                                                                 38516 CGCCCCCCCCCCCCTG 38547
                                                                                                                                                                                                                                                                                   2496 isAlaTrpAspGluGluProLysProLeuLeu 2506
                                                                                                                                                                                                                                                                                                                                                                                                                                          Sorangium cellulosum 68.75 kb contig.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Location/Qualifiers
1. .1826
                                                                                                                                                                                                                                                                                                                                                        AAZ55887 standard; DNA; 68750 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          6251. .21749
                                                                                                                                                                                                                                                                                                                                                                                                       (revised)
(first entry)
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Polyangium cellulosum.
                                                                                                                                                                                                                                                                                                                                                                                                      15-SEP-2003
10-APR-2000
                                                                                                                                                                                                                                                                                                                                                                                AAZ55887;
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qq	36449TACCGGGTGAACCCCGGTGTGTCTTGAAGGACAGCCCGCAGGCTGATGCTGGGG 36501	ė	
ķ	1712 yLeuSerProArgGluSerSerLeuAlaLeuAsnTyrAlaAlaGlyProArgGlylle 1731	Š	
q		9 8	2076 Glubeu
λ̈	1732 IleAspLeuSerGlnValProHisLeuProValLeuValProProThrProGlyThr 1750	: 셤	
DÞ	36562 AGGCCGTTGAGCCCGTTCTCGCCGATGATCAGCCCGCTGGTGCCGCCGGGGTGTG 36621	ò	2096LeuG
λ	ProAlaThrAlaMetAspArgLeuAlaTyrLeuProThrAlaProGlnProPheSerSer	QΩ	37477 CCGCCGI
DP DP		ò	2115 SerSerS
λŏ	ArgHisSerSerSerProLeuSerProGlyGlyProThrHisLeuThrLysProThrThr	qa	37537 CCGGCC
අු	CCGTTGCCGATCAGCACGGCGTTCGGGACC	ò	2135
کن ک	1791 -ThrSerSerGerGluArgGluArgAspArgGluArgAspArgGluArgAspArgGLu-A 1810	q	37594 GGTCGCC
90	GAGCICGAATICCCACCGGIGICAGCGCCAAAACCCGCCGGCCGCCGCCGTGGTGGCGCGCGTGGTGGCGCGCGGTGGT	ζō	2140
λŏ	rgGluLysSerIleLeuThrSerThrThrValGluH18Alarrolle1rpArgF10G	qg	37654 GTGTGC
අු	CCGAACACCCCGGCCGTACCGCCGTCACCGCC	ò	2158 ProLeuf
à i	1830 lyThrGluGInSerSerGlySerSerGlySerSerGlyGlyGlyGlyGlyGlyGlyGlyGlySerSerSerA 1850	q	37714 AAGCCG
gg (GTGCCGCCGCTGCTGCCGATGCCGCTGGAAGCAACCGGTGCCGGCGGCACCGGCCGAAGAAGGAAG	δδ	2178 ArgPro
à i	rgproalassernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernissernisser	đ	37771 CCGCCA
ထူ		δλ	2195
ð í	1870 AlaLeuGlnGlnArgProSerValLeuHisAsnThrGLyMetLySGLy11e11e 1887	Ob	37831 GCGCTCK
a a		ò	2206 Ser
à		QQ	37891 GCCCCA
Q	CCGCCGCTGCCGCCGTTGCCGCCGAAGGTGCCGCCGCTGCTGCTGCCGCCAGCACCGCCAGTG	ờ	2222 IleGlul
ò	ArgProAlaAlaThrPheProProAlaThrHisCysProLeuGlyGlyThrLeuAspGly	qq	37930 CCGGCG
අු		ò	2242 TyrPro
à	ValTyrProThrLeuMetGluProValLeuLeuProLysGluAlaProArgYalAlaArg	q	37990 GCCCG
셤		ò	2262 ProGly
à	ProGluArgProArgAlaAspThrGlyHisAlaPheLeuAlaLysProProAlaArg	qq	38032
g	CCGGICCCGCCAIICGCGCCA	ò	2282 MetVal
à i	1967 SerGlyLeuGluProAlaSerSerProSerIysGlySerGluProArgPro 1983	QQ	38038
a .	AACCCGCGGCCCCCCCCGCCCGCCGAAACAGCCCCAACAGCCCGGAAACAGCCCCAACCGGAAACAGCCCCAACCCGGAAACAGCCCCAACCCGGAAACAGCCCCAACCCGGAAACAGCCCCAACCCGGAAACAGCCCCAACCCGGAAACAGCCCCAACCCGGAAACAGCCCCAACCAACAA	ò	2302 ProGlu
ð,	LeuvalProProvatSerGiyHisAlainrileAlaArginirio	QQ	38041 CCGAAC
셤	GCGGTGCCCCCGGCACCGATAAAGTTTTTGGGAGAGGCGGCCTTGGCCG	ò	2322 ThrGly
ò		qq	38059
සු		oy	2342 LeuGlu
λo	2016 AlaSerAlaSerAspProHisArgGluLysThrGinSerLysProPheSerIleGinGiu 2035	<u>අ</u> .	38098 TTGGAT
e e	GTGTCCAGGCCAAACCCGCTGCCGCCGGTGCCGCCGGAGCCGATCCGATCCGATCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCGATGCCATGCCATGCCATGCCATGCCATGCATG	Š	2362 ProLeu
à	LeuGluLeuArgSerLeuGlyTyrHisGlySerSerTyrSerProGLuGLyAlGluPro	셤	38107 CCGCT
e G	37357		

C----- 38058 sececcecerrececerreceserreceisarea de cos de consece de consecuente de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 17890 de 1789 ACCGCTTCCGCCGGCGCCCCGAAG------GACCCGCCG 37929 ------GAG 38040 ----- ccc 38106 SCCGCGTCGAAGCCCAGCCCGCAGACGCCGCCGCGCG------38031 38037 ----- CCGCCG----- 38037 cadrocodrocococococorroaarccorcocochagraecrococococ 37713 uSerAlaAsnAlaPheAsnProLeuAsnAlaSerAlaSerLeuProAlaAlaMet 2381 ------ATACCTGACAGGATTCCGCCGGCCCCGCCGGATCCGCCGTCT 37476 37536 STOGGCGCCGTCGTCGCCGCCGTTGCCGAACAACCACCGG 37536 ccacce --- resececesse celeces de la contrace de la 1593 CCGGTGAGGGCTTGGGGCTCGTTGATCGCGTTGAGGATTTGTTGCTGCAGG 37653 lLysSerLysLysGlnGluIleAsnLysLysLeuAsnThrHisAsnArgAsnGlu 2301 uTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetProAlaIleThrGly 2321 yLeuMetThrTyrArgSerGlnAlaValGlnGluHisAlaSerThrAsnMetGly 2341 uAlailelleArgLysAlaLeuMetGlyLysTyrAspGlnTrpGluGluSerPro 2361 ---ProSerAspLeuTyrLeuProProPspHisGlyAlaProAla---- 2194 provalSerproproGluGlyMetThrGluproGlyHisSerArgSerAlaVal 2241 yAsnThrSerGlnProProAlaPhePheSerLysLeuThrGluSerAsnSerAla 2281 AspLysSerHisLeuGluGlyGluLeuArgProLysGlnProGlyProValLys 2095 SerProLeuLeuGlnThrAlaProGlyValLysGlyHisGlnArgValValThr 2134 SerGluVallleThrGlnAspTyrThrArgHisHisProGlnGlnLeuSerAla 2157 proAlaproLeuTyrSerPheProGlyAlaSerCysProValleuAspLeuArg 2177 2205 2205 ------proGluProAsnLysThrSerValLeuGlyGlyGlyGluAspGly 2221 bleuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlySerLysSer 2261 ProValSerSerProSerLeuThrHisAspLysGlyLeuProLysHisLeuGlu 2075 GlyGlyGluAlaAlaHisLeuProHisLeuArgProLeuProGluSerGlnPro ||| |----siccingecccarctaraccaccaacccaccagraccacca

à	465
90	32904 GGGCTGGGCCTTGACGACTTCGACGAATGCGATCCGCTCGTGGTGACGGAATCTC 32963
λ	475 LygAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArgArg 490
Db	32964 GAAGATGCGTCGATTCGTTGTTAGCCGGAGGAACGACGCCCACACTAGGTTCGGCACTGT 33023
<i>≿</i> o	GlyLysSerGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnG
q	
λ̈́O	
qq	33084 ATGGTCACCGCGGTGTGCAGCCAGCAGCACGTTGACGTCGATGAGGAACATCGCCTATTT 33143
à	
QQ	33144 GIGCCIGICCAGGCTCACTICCGCGAGITCAGITCCAGACCCICGICGAGCACTICGGAC 33203
Š	545 LeuLeuLysGluLysThrAspAspThrSerGlyGluAspAsnAspGluLysGluAlaVal 564
qq	33204 AACACCGTATTCGAGGTTAGGTCGATACCTGGCCGGACCGGTCCGGCGTCAAAAACG 33263
ò	565 AlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgArgLysGly 580
q	33264 GGGACGGTTGGCCGGCGCCGCCGGTACGGCGGCGGCGGCGGCCGCCGAAGGGCGTCT 33323
à	laAsnGlu
셤	33324 TCGATCACAGCGCCCAGCGATTAACCACGCTCGCGGGCCCGGCGTTTGGCGGTAGCCAGT 33383
à	591 AsnSerGluGlu 594
gg.	33384 AGTICATCCGAGATTGACACGGTGGGTGCGCATGATGCTCAGGATAGCGCATCTACGGCAT 33443
à	595
d d	33444 CATCTGCGGTGAGCAACTGATGCCCTCAACGCCGCGTGGTGGTCGCAGGTCTGCCTGTAT 33503
λŏ	607 tGluLeuAsnGluSerSerArgTrpThrGluGluGluMetGluThrAlaLysLysGlyLe 627
QQ	decerraagreegrereeceageageage
È	627 uLeuGluHisGlyArgAsnTrpSerAlalleAlaArgMetValGlySerLysThrValSe 647
QQ	33547 -TGCCGCCGGCACCGGCGTGGCCCCGGCTTGCCGG33582
à	SysLysAsnPheTyrPhe
Q	33583 -CGTTGCCGCCGTTGCCGCCGTTGCCGATCAGCACGCGTTGCCGCCG 33629
ờ	nGlnHisLysLeuLysMetGluLysGluArgAsnAlaArgArgLysLysLysLysBLysBlaPr
g Q	
λ̈́σ	687 oAlaAlaSerGluGluAlaAlaPheProValValGluAspGluGluMetGluAl 707
g G	
à	
qq	33715TCACCGCCGGTGCCGCCGGTACCGGCGCTAGAGCGTT 33752
à	727 rGlyAsnGluValProArgGlyGluCysSerGlyProAlaThrValAsnAsnSerAs 747
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ò	747 pThrGluSerIleProSerProHisThrGluAlaAlaLysAspThrGlyGlnAsnGlyPr 767
G G	33801 GCCGCCGTTCCCGCCGCGCGCCCG

767	oLysproproAlaThrLeuGlyAlaAspGlyProProProGlyProProThrProPro- 7	86 3884
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78	argarghn:SerargalaProlleGluBroinFroAlaserGluatainica o	33944
20 0		. 4
33945	YALBETOILLETOILLETOILLONGE GAAGCTGGCGCCACCGCTGCCGCCGGCGCCACGGAGGCGCCGAGGAGAGTAGGCCGGC 3	34004
		832
34005	GTTGCCGCCGCCGCCGCCGCCGCCGCGGTGTTGTTCCCGGGCCCTCCCGGGGCC	34055
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34056	gecesceccecestrece-eccestrecestrecescerreces	34114
847	aAlaGluGluLeuAlaValAspThrGlyLysAlaGluGluPr 8	861
34115		34174
861		879
34175		34234
880		893
34235		34294
893		606
34295	C	34339
606	rGlyAlaproGlnAspSerAspSerSerAlaThrCysSerAlaAspGluVa	926
34340	cecceccccareccaccaccccrareaagrrcccacccccccccc	34394
926	lAspGluAlaGluGlyGlyAspLysAsnArgLeuLeuSerProArgProSerLeuTh	946
34395		34415
946	rProThrGlyAspProArgAlaAsnAlaSerProGlnLysProLeuAspLeuLysGlnLe	996
34416	GAAGACCCCACCGGCGCCGCCGG-CCCCGGCCGATCCCTCCAGCACCGGTGGCT	34467
996	uLysGlnArgAlaAlaAlaIleProProIleGlnValThrLysValHisGluProProAr	986
34468	ATCCGGCGGGTCCCG-CCATTGCCACGGTACCGAACAAGATCCGGCGGC	34517
986	gGluAspAlaAlaPro	66
34518	cccccccdccccccccccccccccccccccccccccccc	34577
666	oproproglnAsnLeuGlnProGluSerAspAlaProGlnGlnProGlySe	1017
34578		34637
1017	rSerProArgGlyLysSerArgSerProAlaProProAlaAspLysGluAlaPheAlaAl	1037
34638	GGCGCCGTTGCCGAACAGCCACCCGCCGGCCCCGCCG	34674
1037	aGlualaGlnLysLeuProGlyAspProProCysTrpThrSerGlyLeuProPh	1055
34675	TCAGCCCCGGTTCCAGGAGTCCCGTTGGCGCCGTTGCCGATCAGCGGGCGG	34730
1055	eProVal	1057
34731	GAGCGTCTGGAAGGGCTCGTTCACCACATTGAGCACATTTTGCTGCAGGGTGTGCAGTGG	34790

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1.42 Protection Control	65625 CGAATACGC	1726 AlaGlyProArgGlyIle 65575 GATAACCCTCGAAC	1746 ProThrPro		1749GlyThrProAla	65497 CGGTCCGGCTCCCCCGCA	1767 ProPheSerSerArgHis	65437 CGACTTCCAGCAGCAACC	1784HisLeu 	65377 CGAAGAAACCGGCATCGA	1800 ArgAspArgGluArgAsp 	obsil AdditionAndreGAI	1820 InrvalgiumisalaPro 65284 ACCCGCCTGATCCCAGC	1840 SerSerGlyGlyGlyGly	65224 CCACCA	1860 HisSerProlleSerPro	65176 GACAGCCCATCCCCA	1880 AsnThrGlvMetLvsGlv	65131 CCGCCACAGCCGCCG	1900 SerThrSerThrSerSer	65074 GCAAGAACCGAACG	1918 HisCvsProLeuGlyGlv	65014 GAACGCCCA	1938 LeuProLysGluAlaPro	65005GCACCC	1958 AlaPheteuAlaLysPro	64957	1978 GlySerGluProArgPro	64939 CACGATCGGCCCCGACCT	1998 ProAlaLysAsnLeuAla	64903 CCAGCACCG	2012ProAlaProProAla	64852 GATCCAGGTCCGGCAGCC	2031 PheSerIleGlnGluLeu	64793 TTGACC
	q _Q	දි දි	ò	7 A	ò	QO	ò	qq	ð.	ପ୍ର	& €	<u>a</u> 2	Š €	8 8	; A	3 8	G 42	δ	; 셤	ò	: 셤	8	3 A	ò	q	ò	đ	ò	셤	λō	qq	8	q _Q	ò	đ
6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6			66543 Agggcrdadcgcrgccddrgrrrcrcccacacgarcagaacddrggaaggrccac	1449 GlySerIleThrGlnGlyThrProLeuLysTyrAsp	66483 GGTCGATTCCGAATCGGGACCGCTCTTGTCCCGCAGTCTCGTGTTTTTCGAGGCTTG	1461	66423 IICGAGGAIGACGIGIGGGIIGGCGCIGAICCCGAACGACGACGACGACGCCCGCC	66363 AGGANGGCTGTTGCCGGGGGCTGTTCT-CCGTGAGGAGCTGGACATCCCCCGCCG	1487 'ValHisProLeuAspValMetAlaAspAlaArgAlaLeuGluArgAla		1503CysTyrGluGlu	66244 GCATCGCCATCACCATCTTGATCACCAGCGACACCCGCCGCCGCCGCCTGCGTATGACCAA	1507SerLeuLysSerArgProGlyThrAlaSerSerSerGlyGlyGerlleAlaArgGly	66184 TATTCGACTTCACCGACCCCAACCACGACGGCCGATCGCTATCCCGCCCTGCCCGT	1526 AlaProValIleValProGluLeuGlyLysProArgGlnSerProLeuThrTyrGluAsp	66127CAGTGGCAATCAGAGCCTGCGATCGGAT	1546 HisGlyAlaProPheAlaGlyHisLeuProArgGlySerProValThr	66094 caccaadcerggriccagreechigegeerecaegegreecheareagaeaegaaaae	1562 MetArgGluProThrProArgLeuGlnGluGlySerLeuSerSerSerLysAlaSerGln	66034 CAGCACTCGCCAACGCCTGCGTGATCACCCGCTGCTGCGAGGGCCCATTCG	1582 AspArgLysLeuThrSerThrProArgGluileAlaLysSerProHisSerThrValPro	65983GGCCGTCAAACCATTCGACG	1602 GluHisHisProHisProlleSerProTyrGlUHisLeuLeuArgGly :::	65962CACCATCCTGATTCACCGCGCTACCCCGCACGACCGCAGGACCGCGTGCCCGTTGC	1618 ValSerGlyValAspLeuTyrArgSerHisIleProLeuAlaPheAspProThrSerIle	65905 GCCGGGCATCCGACAGCCGCTCCAACACACCGCGCCTTCT	1638 ProArgGiyileProLeuAspAlaAlaAlaAlahiyriyrLeuFroArgHisLeuAlaPro	65858 CCCCAACCGGIGCCAICCGCAGCCGCCGCAAACGACIIGCACCGCCCAICCGCCGCCGCCGCCGCCGCCGCCGCCGCCGCCGCCGCCG	1658 ABBEROHISLERIYEKTO	03001 AMGCCCCG11GACGCGAAAACICCACGAACAICCCGGGGGGGGGG			65681 ACCAACGACGA-CGAACACGCCGTATCCACCGTCACCGCAGGACCCTCGAAAACCAAA	1706 ArgAlaAspMetLeuArgGlyLeuSerProArgGluSerSerLeuAlaLeuAsnTyrAla	

65625	CGAATACGCAACCCGACCCGAAAACACGCCTCCCGCGCTGC-CATTACCTA 6	5576
1726	AlaGlyProArgGlyIleIleAspLeuSerGlnValProHisLeuProValLeuValPro	745
65575	GATAACCCTCGAAACCCT 6	5558
1746	ProthrPro	748
65557	CCGGCGCCCTCGTGATGAACCGCGCGCGTAGTCGTGGTGCTATCAGCCCCGCGAACACGC 6	5498
1749	GlyThrProAlaThrAlaMetAspArgLeuAlaTyrLeuProThrAlaProGln	1766 65438
76	ProPheSerSerArgHisSerSerSerProLeuSerProGlyGlyProThr	1783
65437	caacticcagcagcagcagcagcagcagcagcagcagcagagaaatcc 6	5378
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65377	CGAAGAAACCGGCATCGAGCATCTCGCAGGAAGCCACCTCACACACA	5318
1800	ArgAspArgGluArgAspArgAspArgGluArgGluLysSerlleLeuThrSerThrThr 1	819 5285
1820	ThrvalGluHisAlaProlleTrpArgProGlyThrGluGlnSerSerGlySerSerGly 1	839
65284	 ACCOGCCTGATCCCAGGATCGTCCGGGAACTCCGCCACCGCATCCAAGCCCCCGG 6	5225
. 1840	SerSerGlyGlyGlyGlyGlySerSerSerArgProAlaSerHisSerHisAlaHisGln 1	859
65224	CCACCAAACGCCAAAGCTCCTCCGGCGAGGACACCCCCACCCGGATAAC 6	5177
1860	HisSerProlleSerProArgThrGlnAspAlaLeuGlnGlnArgProSerValLeuHis 1	679
65176	GACAGCCCA TCCCCACGATCACAACGGGATCGTCAGCAGTCACCG 6	5132
1880	AsnThrGlyMetLysGlyIleIleThrAlaValGluProSerLysProThrValLeuArg 1 :::	99
65131	9	S.
1900	SerThrSerThrSerProValArgProAlaAlaThrPheProProAlaThr 1 ::: ::: :::	917 5015
1918	HisCysProLeuGlyGlyThrLeuAspGlyValTyrProThrLeuMetGluProValLeu 1	937
65014	GAACGCCCA	9009
1938	LeuProLysGluAlaProArgValAlaArgProGluArgProArgAlaAspThrClyHis 1	957
9 6		, ,
1958 64957	ALaPheLeuAlaLysProbroAlaArgSerGlyLeuGluProAlaSerSerProserty 1	4940
1978		997
64939	CACGATCGGCCCCGACCTCGACCGCACCCGAPTGCC 6	4904
1998	ProAlaLysAsnLeuAlaProHisHisAlaSerProAspPro	011
64903		4853
2012	ProAlaProProAlaSerAlaSerAspProHisArgGluLysThrGlnSerLysPro 2	030
2 6	The control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the co	0.50
79	TIGATOGCGCTCCGCCGCGCGCTGGCTGCACGTCGCGCAACAAC 6	· -

1110		67527 GACGCCGTCC			1126 erGlnGlyMe	67414	1145	67392 GCGTGCGAT	1160 ysLeuAlaP	 67335 ATGCGAAAC	1177 lnAlaGlyP	:: 67275 GTCAAGCAC	1197 lyThrAlaL	 67227 G-ACCGCG-	1217 alProSerA	 67177 GCTGCTCCT	1230 hrHisGlyT	67117 ACCGGGCAC	1250 spserPros	::: ::: 67075 ACAACAC-T	1270 yrGluglyL	:: : 67017 ACCC-GGCC	1290Ser	::: 66967 ACGCGAACA	1302 isGluThr	:: : 66907 ACACCACAI	1322 leSerSer ²	66847 CGTCAAACC	1339	66788 CACTGTCCC	1349 isHisIle	66728 ACCAC	1369 spTyrLeu	66695 GCTTCCAA	1389 roSerArg	66636 CAACGCGG	1409 ro-AlaHi
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67527	GACGCCGTCGGCGAAGGGGACGGGCTCTGGACCTGACGCACCCAATACTCCGGCGTAGC 67468
1113	
67467	CATCACACCTGCGGTGTCGAGCTCACCGGTCACGTCGACACGACGGTAGGT 67415
1126	GlyMetSerValGlnLeuHisValProTyrSerGluHisAlaLysAlaPro 1144
67414	673
1145	160
67392	673
0	77
67335	9
1177	InalaGlyProProGluSerLeuGlyValProThrAlaGlnGluAlaSerValLeuArgG 1197
2 1	roserthraray 121
67227	
ı	alproSerAspSerAlaileThrTyrArgGlySerileT 1230
17	
1230	hrHisGlyThrProalaAspValLeuTyrLysGlyThrIleThrArglleIleGlyGluA 1250
67117	ACCGGGCACGACCGCCACCATCCGAGCCGCATCCGGAACG 67076
1250	spSerProSerArgLeuAspArgGlyArgGluAspSerLeuProLysGlyHisVallleT 1270
67075	670
1270	289
67017	0
59	SerLygGluAspGlyArgSerSerSerGlyProProH 1302
66967	
1302	7 7 7
66907	9 6
1322	338
66847	667
1339	349
66788	CACTGTCCGCCTGACCGGCGAACACGAACCCCGCCAGAGACCGGGGATCCCGAC 66729
1349	isHisIleArgGlySerIleThrGlnGlyIleProArgSerTyrValGluAlaGlnGluA 1369
66728	
1369	spTyrLeuArgArgGluAlaLysLeuLeuLysArgGluGlyThrProProProProP 1389
66999	GCTTCCAACCCGGACAACAACGTTCGCGGTCCGCTCCCA-GCACCACCGCCCCCTCCTC 66637
1389	rArgAspLeuThrGluAlaTyrLysThrGlnAlaLeuGlyProLeuLysLeuLysP 1409
66636	GGGGCCCGGGAGGAAGCCAACGAAAACG
1409	ro-AlaHisGluGlyLeuValAlaThrValLysGluAlaGlyArgSerIleHisGluile 1428

Db 69634 CCACGCCACAACGGAAATTCCGCC Qy 581 rg11eThrArgSerMetAlaAsnGlu	69454 CACCGCGTTCGG 619 luMetGluThrA 69394 AATCAGCACGCA 633 - ABNTTPSerA 69338 GCGACCGAAGCC 652 heTyrPheAsnT		Oy 727	8 8 8 8
	Qy 284 PheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPheCysGlnArgTyrAsp 303 Db 70461 GACGGGCCCTGGTCGTCGCGCTGGCGTTGGGTTCGTAGCAT 70417 Qy 304 GlnLeuMetGluAlaLeuGluLysLysValGGTLArgIleGluAsnAsnProArgArgArg 323 Db 70416 CTCGTCTGCCAGTTCCTGGTGCAGGAACCGGCCCTCGGCGTCGG 70366 Qy 324 AlaLysGluSerLysValArgGluTyrTyrGluLysGlnPheProGluIleArgLys 342 Db 70365 ATAGTCGAACCCGCAGCACCCCCAGCACCCCGGCAAGGCGGTTCCG 70306 Qy 343 GlnArgGluLeuGlnGluArgMetGlnSerArgValGlyGlnArgGlySerGlyLeuSer 362	Db 70305 CAACTCCACACCACAACGAATCAAAACCCAGATCCCGGAAAGCGCGCTCGACCCCGAC 70246 363 MetSeralaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSerGluGln 382 :::	Qy 422 ITyrLysAspArgGlnValMetAsnMetTrpSerGluGlnGluLysGluThrPheArgGl 442 Db 70086 CTGCTCGGCCAGGAAAGTCCTGTCCAACTTCGCCGGAACTACCGCTGC 70039 Qy 442 uLysPheMetGlnHisPro	Qy 472 uThrLyslysAsnGluAsnTyrLysSerIeuValArgArgSerTyrArgArgArgGlyLy 492 Db 69934 cCGTCATCCCGCTCGCTCGCCCAGGCCCAGCCCAGCCAGGCAGTCCC Qy 492 sSerGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnG
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69634	CCACGCCGCACAACGGAAATTCCGCCGGGACCGCCAGCAACGACTCCAGATCAGCAC 69575
581	593
69574	GATCCGCGACGTCGCACCACGTCGACCACGGCACCGAGGCCGTTCAGCTCACCCA 69515
593	
69514	
613	SerArgTrpThrGluGluG 619
69454	
619	luMetGluThrAlaLysLysGlyLeuLeuGluHisGlyArg632
69394	AAATCAGCACGGTTCGGTC-GGGATCCGCCACAACCGCGGGCACGTCGATCTGCCCG
633	ABNTTPSerAlaIleAlaArgMetValGlySerLygThrValSerGlnCygLygAsnP 652 :::
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2 (1	GCCAGCACAAACTGCCCCGGGTTCTCCACCTGCGCGGCACGCAACAACCCCCCAAACGGC 69
687	ProAlaAlaAlaSerGluGluAlaAlaPheProProValValGluAspGluGluMetGlu 706
69161	
707 69116	AlaSerGlyValSerGlyAsnGluGluMetValGluGluAlaGluAlaLeuHisAla 726
727	
95069	TGGAGCAGAGCCAACGTCTGCTGCGAAGTGCGGCTGCGCCGGTGCCGGATCC 68997
735	GlucysSerGlyProAlaThrValAsnAsnSerSerAspThrGluSerIleProSerPro 754
755	
68939	
763	
68819	
774	yAlaAspGlyProProProBlyProProProArgArgArgThrSerArgAl 792
79	aProlleGluProThrPro
68759	GCCAGCTGTCCCGACGACACGCTTCGTAGCCTGAGAATCGATCG
808	aprothrProProProProAlaProPro
818	oProProValValProLysGluGluLysGluGluGluThrAlaAlaAlaProProValGl 838
838	uGluGlyGluGluGlnLysProProAlaAlaGluGluLeuAlaValAspThrGlyLysAl 858
68595	TAAGAACGGCACCACGCGCCGCCTTGCTCCGG 68

FT misc_feature 4775348310 FT //tag= ar //note= "Dehydratase domain (DH5): part of extender module FT FT misc_feature 4922649771 FT misc_feature //tag= as //note= "Beta-ketoreductase domain (KR5): part of extender FT module 5" module 5" //stag= as //note= "Beta-ketoreductase domain (KR5): part of extender FT misc_feature 5000950254	Alignment Scores: 3.3e-10 Length: 80161 Pred. No.: 631.50 Matches: 680 Score: 31.104 Conservative: 317 Best Local Similarity: 21.894 Mismatches: 958 Ouery Match: 2 Gaps: 139	AAZ21501 (1-80161) Propro	Db 71206 ACAACACCAAACCAGCACCCTAACCCAAACCGAAACGGAACCCTAACCCAAACCGAAACGGAACCGAACCGAAACGGAAACGGAAACGGAAACGGAAACGGAAACGGAAACGGAAACGGAAACGGAAAAAA	71109 91 71079	Db 71019 CAPATGCAACGACGACGACGAACACGCAGTATCCACCGCACGCA	132 70854 152 70806	171 70746 191 70716	Oy 211 GlubysProValSerProProProlleGluSerbysHisArgSerbeuValGlnIle 229
module 2" 3062931621 /*tag= aa /note= "Acyl transferase domain (AT2): part of extender module 2" 3169732254 /*tag= ab /*tag= ab /*note= "Dehydratase domain (DH2): part of extender module 2"		3488630295 3488630295 //tag= about arrier protein domain (ACP2): part of extender module 2" 3541944331 /*tag= af /product= "spnc" //tote= "Spinosyn biosynthesis protein C"		U	ψ.	= 4 = 4	/*tag= an /*tag= an /*tag= an /*tag= an /*tag= an //note= "Acyl carrier protein domain (ACP4): part of extender module 4" 4496659752 /*tag= appn" /*tag= an //note= "Spinosyn biosynthesis protein D" //note= "Spinosyn biosynthesis protein D" /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*tag= appn /*t	= rV · = rV
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extender module 1"
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21126. .22379
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note= "Spinosyn biosynthesis protein O"
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complement(18541, .19713)
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16556. .17743
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/note= "Spinosyn biosynthesis
20168. .20995
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 21758 CIGTCGTTGGAAGG-----CATACGTTGGCAACTCGAGCTGCCTAGCGCCGGTACCGGC 21705
                                                                                                                                 ccgaaccagreccecca-----GCGTTCCGTCCGGCCGAGCTCGACGACCGTGCC 21510
                                                                                                                                                                                                                    2403 laLygValSerGlyArgProSerSerArgLygAlaLysSerProAlaProGlyLeuAlaS 2423
                                                                er----GlyAspArgProProSerValSerSerValHisSerGluGlyAspCysAsnA 2441
                                                                                                             2441 rgArgThrProLeuThrAsnArgValTrpGluAspArgProSerSerAlaGlySerThrP 2461
                                                                                                                                                         2461 roPheProTyrAsnProLeuIleMetArgLeuGlnAlaGlyValMetAlaSerPro--- 2479
                                                                                                                                                                                                    2480 ------ProProProGlyLeuProAlaGlySerGlyProLeuAlaGlyProH 2495
                                                                                                                                                                                                                                                                                                                                                                                     DNA fragment of Saccharopolyspora spinosa containing biosynthetic genes.
                                                                                   21644 CACCGAACGGGTCTCGTCCCCTGCGCATCAGTGGGATCG-------
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         notes "Protein involved in transcription control" 2024. .2791
                                                                                                                                                                                                                                                                                                                                                                                                         Spinosyn biosynthetic enzyme; open reading frame; ORF; insecticidal microlides; arachnid; nematode; insect; polyketide; polyketide synthase; PKS; extender module; initiator module; acyl transferase domain; AT; acyl carrier protein; ACP; beta-ketosynthase domain; KS; KR; dehydratase domain; DH; enoyl reductase domain; ER; beta-ketoreductase; insecticide; ss.
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                                                                                                                                                                                                                                                2495 isHisAlaTrpAspGluGluProLysProLeuLeuCysSer 2508
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083. 8450
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complement(3416. .4165)
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product= "ORFL16"
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AAZ21501 standard; DNA; 80161 BP
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product= "SpnO"
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apheAsnProLeuAsnAlaSerAlaSer-LeuProAla-----AlaMetProIleThrA 2385 |::: |||||| :::::: GATTCGGTACCGGCAGGAATCGACGAGCGGTTGCGGAGTCCGCTTCGCCAGGAGGA 21936 caacecedehanacarecereaaegargegeargregregregregeeecaagareead 21876 decedecalectacetreceagaresectredereaacesegereeag---aactetrese 21819 | | | | ::: | | | | recarcentregaeadeacearceaaceace 22116 glysaljaLeuMetGlyLysTyrAspGlnTrpGluGluSerProProLeuSerAlaAsnAl 2367 5 laalaaspGlyargSerAspHisThrLeuThrSerProGlyGly------GlyGlyLysA 2403 AAAAGGGGCTTCGGCAAGGCCCGCAGAAGCTGCACAGTCAGCGGAA 22284 |||::: |ACCAGTT------GGCAAGCAGAGAATCCCGGAGCAGGCGATTCATCGAGCCC 22336 uMetThrTyrArgSerGlnAlaValGlnGluHisAlaSerThr-------2338 : gecaggicgaicaagccaccccagaacicgggigggiggiggiggiggiggiggiggigg | :::||||||| T-----TCCAACGCGACGCGCGCGCGCGCCACCGCCA 22344 ------cchacacacceccichaccegecacccaatcdgicgtcccagtccdat 22475 oGlyAsnThrSerGlnFroProAlaPhePheSerLysLeuThrGluSerAsnSerAlaMe 2282 tValLysSerLysLys-----GlnGluIleAsnLysLysLeuAsnThrHisAsnArgAs 2300 ...----be 2324 cccgaagcgrccgcrccrccacaggcrcaacaccacgcrccagcaccggcc 22685 ||||::: | |||-::: | GCACGAACACCACCACGGCGAATGGCGACTTGAT-----CCTCGCCACCATCCG-- 22522 uProdiyHisSerArgSerAla-----valTy 2242 accardededeaseaceceádaceaseceaseaceaseaceasedadedadedarrecaades 22574 lleudspleudrgdrgProproSerAspleuTyrLeuProProProAspHisGlyAlaPr 2193 oAlaArgGlySerProHisSerGluGlyGlyLysArgSerProGluProAsnLysThrSe 2213 valLeuGlyGlyGlyGluAspGlyIleGluProValSerProProGluGlyMetThrGl 2233 sLeuGlyGlyGluAlaAlaHisLeuProHisLeuArgProLeuProGluSerGlnProSe 2115 rSerSerProLeuGlnThrAlaProGlyValLyBGlyHisGlnArgValValThrLe 2135 uAlaGln-----HislleSerGluVallleThrGlnAspTyrThrArgHisHisProGl 2153 nglnLeuSerAlaproLeuProAlaproLeuTyrSerPheProGlyAlaSerCysProVa 2173 -----IleAr nGluproGluTyrAsnIle-----SerGlnProGlyTh rproLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlySerLysSerPr rGlullePheAsnMetProAlaileThrGlyThrGly-------AsnMetGlyLeuGluAlaIle------

1604 24797 1617 24737 1627		1663 24500 1664 24440	Qy 1671 gGlyTyrProAspThrAlaAlaLeu Db 24386 AGGCCTCCCAGGACACCACCAGCA Qy 1681	1683 24266 1701 24206	1703 24146 1720 24086	Dy 1/36 MYALFDO	Db 23906 CTCGAATATCTCGTCGTCGCCCCC Qy 1784 SLeuThrLysProThrThrThrSer Db 23846 TACTTCAGCTGCGTGTTCCAC Qy 1804 GASPArgAspArgGluArgGluLys Db 23789 GTCCGTGCCGGGCTTCGCCGGTG
1375		1454 yThr	1469 BLyBHiBABDVal	1480 oGlyArgThrPhePr 1485 15301 GGCAAGGCGACCACTGTCCGGCCAGGCGTGTTCTCCGTCGAGAAGCCGGAACCC 25242 1485 oProValHisProLeuAspValMetAlaAspAlaArgAlaLeuGluArgAlaCysTyrG1 1505	udluserLeuLysserArgProdly	1522 eAlaArgGlyAlaProValIleValProGluLeuGlyLy8ProArgGlnSerProLeuTh 1542 25064 CCGTACGTGG	ACGACAGCCCCGCACGCCAACGCCTGGGTGATCACCCGGCTGCTGCGGCGACCATC rSerLysAlaSerGlnAspArglysLeuThrSerThrProArgGluIleAlaLysSerPr :::::::::::::::::::::::::::::::::::
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ò		q	26272 CATGCGACACCGCAACCACC
qq		λö	1251 SerProSerArgLeuAspArgGlyA
δ		qq	26242 ACCCCTGATCGTGCAACCGATCAGG
QQ		ò	1271 GluGlyLysLysGlyHisValLeuS
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đ	27066 GGATTCCCCGGGCGGTCCCACCAGCACCTGAACCTGGACCGCTCCATG 27019	δ	1291 LysGluAspGlyArgSerSerG
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qq	26970 ACAACCGAGGCGTCGCCGACGTGCCACCACCACCACGATCGCGGTGCCGGGGACGAC 26911	λ	1312 AspMetMetGluGlyArgValGlyA
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----GlnHisHisHislleargGlySerIleThrGlnGl 1358 |||||||||||||||||| rgccccaACACCACATCCGG------CCGAA 25943 ||||||| GGTCCCGCACCCCATCCAGCAGAGGATCGACCTGAA 26108 GCGGCCTCCGACAACACACACAGCGGCGATGCA 25991 ATGGCACCAGCAGGGAAGGCTGCATCAACC--- 26051 CCAGCATCGGCT---CCATATGGGGCGAATGGAAAG 26273 ------GluarghisSerProhis-Hi 1344 lagingluAspTyri,euArgArgGju----- 1374 :::||| |||||| -------ATCGGCAGTTCCGGTGCGTGAT 26339 yArgAlaIleSerSerAlaSerIleGluGlyLeuMet 1331 ArgGluAspSerLeuProLysGlyHisVallleTyr 1270 1290 uSerTyrGluGlyGlyMetSerValThrGlnCysSer rGlyProPro---- 1301 -----HisGluThralaAlaProLysArgThrTyr 1311 ctracceatcaccedearccceccacccccccccccs 26554 AlaLeuGlySerValProGlyGlySerIleThrLys 1210 SerAspseralaileThrTyrArgGlySerlleThr 1230 ıTyrLysGlyThrIleThrArgIleIleĞlyGluAsp 1250 luHisAlaLysAlaProvalGlyProvalThrMetG 1151 ysLysLeuAlaProPheSerGlyValLysGlnGluG 1171 lyProProGluSerLeuGly-ValProThrAlaGln 1190 -- ProThrileSerAsnProProProLeuIleSerS 1111 luArgGlnIleGlyAlaIleSerGlnGlyMetSerV 1131 ro---LeuProLeuGjyLeuHisAspThrAjaArg- 1093 ---¢Tcgggaga------

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ò	174GlnasnMetAspArgValAspArgGluIleThrMetValGluGlnGlnIleSerLy 192	3 8	_
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641	ValGlySerLysThrValSerGlnCysLysAsnPheTyrP 654
28108	ACGCCTGCCACGACTCAGGTGTGCCGTCCACCAGGACGAAGCGACCCGGGTTCT 28049
654	heAsnTyrLysLysArgGlnAsnLeuAspGluIlsLeuGlnGlnHisLysLeuLysMetG 674
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692	luGluAlaAlaPheProProValValGluAspGluGluMetGluAlaSerGlyValSerG 712
27928	GCTCGCAGCCACGACTGCACGAGAGCCAGCAGATCGGAGGTCGACTCTTGGGCGC 27869
712	1948mGluGluGluMetValGluGluAlaGluAlaLeuHisAlaSerGlyAsmGluValP 732
732	roArgGlyGluCysSerGlyProAlaThrValAsnAsnSerSerAspThrGluSerIleP 752
27808	CCGGGGCAGGAACGTCCAAGGCCGCACGGAACTCATCCAGGGTCGCGAAC 27753
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AAF88316; 28-AUG-2001 (first entry) S. spinosa DNA fragment SEQ ID 5. Forosamine; trimethylrhamnose; polyketide synthase; biosynthesis; spinosyn; polyketide aglycone; transgenic plant; insect resistance; macrolide; insecticidal; ds. Saccharopolyspora spinosa. DE19957268-A1. OS-NAR-2001. 29-NOV-1999; 99DE-01057268. C9-NOV-1999; 99DE-01057268. C7-AUG-1999; 99DE-01040596. CPARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. Db WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp; German.		qq
S. spinosa DNA fragment SEQ ID 5. S. spinosa DNA fragment SEQ ID 5. Forosamine; trimethylrhamnose; polyketide synthase; biosynthesis; spinosyn; polyketide aglycone; transgenic plant; insect resistance; macrolide; insecticidal; ds. Saccharopolyspora spinosa. DE19957268-A1. 08-WAR-2001. 29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (PARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; Eberz G, Moehrle V, Froede R, Velten R, Salas JA; Eberz G, Moehrle V, Froede R, Velten R, Salas JA; Ecombinant production of insecticidal spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp; German.		ò
S. spinosa DNA fragment SEQ ID 5. Forosamine; trimethylrhamnose; polyketide synthase; biosynthesis; spinosyn; polyketide aglycone; transgenic plant; insect resistance; macrolide; insecticidal; ds. Saccharopolyspora spinosa. DE19957268-A1. 08-MAR-2001. 29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (FARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyn biosynthese. Claim 7; Page 74-91; 354pp; German.		q
Forosamine; trimethylrhamnose; polyketide synthase; biosynthesis; spinosyn; polyketide adlycone; transgenic plant; insect resistance; macrolide; insecticidal; ds. Saccharopolyspora spinosa. DE19957268-A1. 08-WAR-2001. 29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (FARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; Eberz G, Moehrle V, Froede R, Velten R, Salas JA; Everandinant production of insecticidal spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp; German.		ζ
macrolide; insecticidal; ds. Saccharopolyspora spinosa. DE19957268-A1. 08-MAR-2001. 29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (FARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Oy Claim 7; Page 74-91; 354pp, German.	iosynthesis; ct resistance;	đ
Saccharopolyspora spinosa. DE19957268-A1. 08-WAR-2001. 29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (FARB) BAYER AG. (FARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp, German.		ò
DB19957268-A1. 08-MAR-2001. 29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (FARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Qy Claim 7; Page 74-91; 354pp, German. Qy		qq
08-MAR-2001. 29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (FARB) BAXER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Db Claim 7; Page 74-91; 354pp, German. Oy		ò
29-NOV-1999; 99DE-01057268. 27-AUG-1999; 99DE-01040596. (FARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp; German.		q
27-AUG-1999; 99DE-01040596. (FARB) BAXER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp, German.		ò
(FARB) BAYER AG. Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp, German.		<u>අ</u>
Eberz G, Moehrle V, Froede R, Velten R, Salas JA; WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp, German.		ò
WPI; 2001-267102/28. New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7; Page 74-91; 354pp, German.	,	q
New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives. Claim 7, Page 74-91; 354pp, German.		λõ
Claim 7; Page 74-91; 354pp; German.	thesis, useful for their derivatives.	q
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is invention describes a novel method nucleic acid (I) and its encoded ilypeptide (II) containing at least one region that encodes an enzymatic trivity involved in biosynthesis of spinosyns. (I) are used (i) to lentify, inactivate or modulate genes involved in the biosynthesis of lentify, inactivate or modulate genes involved in the biosynthesis of lentify, inactivate or modulate of polyketide synthases; (ii) for il); (ii) to generate a library of polyketide synthases; (iii) for liding forosamine or trimethylthamnose to a spinosyn or polyketide lycone; and (iv) for recombinant production of the corresponding lycone; and (iv) for recombinant production of (II), their precursors or rivatives, including production of transgenic plants that express (II) are takers for sequencing of the Saccharopolyspora spinosa genome. (II) are croides with insecticidal, but not antibacterial, activity, and can so be used to raise specific antibodies, useful for identifying so be used to raise specific antibodies, useful for identifying pression clones in a gene bank. Cells transformed with (I) may produce I) at significantly increased levels or produce new derivatives of I). This sequence represents a genomic DNA fragment of the S. spinosa nome which contains the coding regions for proteins involved in resamine and trimethylrhamnose biosynthesis
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	LeuserAlaProleuProAlaProleu	GACCCCGGGGGCTCCAGCCGCTCGCC roGluProAsn	2211ValleuGlydly 2217	ArgserAlaValTyrProLeuLeuTyrArgAap	2255 SerArgMetGlySer 2259 38446 AACGCAACAACACCACGACGCATGCACCAGGTCGGAAACACCCTCGGCGGACCTG 38387 2260 LysSerProGlyAsnThrSerGlnProProAlaPhePheSerLysLeuThrGlu 2277 38386 TCGACGCCACCACGTCACAACCACCAGGCGAGATCCGGCGCAATCGCGCCGCCA 38327	2278SerAsnSerAlaMetValLysSerLysLysGlnGluIle 2290	2311 ThrGlullePheAsnMetProAlaileThrGlyThrGlyLeuMetThrTyrArgSerGln 2330 38218 GCGGGTCAGCCAACCCGGTCAGGTGG-AGTTCCGTGCCAAGTGCC 38169 2331 AlaValGlnGluHisAlaSerThrAsnMet
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1784HisLeuThrLysProThrThrSerSerSerGluArgGluArgAsp	99853 CCACCAAACGCCCAAAGCICCICCGGCGGAGGACACCCCGGCCGGATAAC 59806 1860 HisserProlleSerProArgThrGlnAspAlaLeuGlnGlnArgProSerValLeuHis 1879 19805 GACAGCCCATCCCCACGATCAACGGGATCGTCAGCAGTCACGG	Oy 1900 SerThrSerTerProValArgProAlaAlaThrPheProProAlaThr 1917	1938 LeuProLysGluAlaProArgValAlaArgPro	1978 GlySerGluProArgProLeuValProProValSerGlyHisAlaThrIleAlaArgThr 19568 CACGATCGCCCGACCTCGACCGCACCC 1998 ProAlaLysAsnLeuAlaPrOHisHisAlaSerProAspPro	2012ProAlaProProAlaSerAlaSerAspErcHisArgGluLysThrGlnSerLysPro	2051 GluGlyValGluProValSerProValSerSerProSerLeuThrHisAspLysGlyLeu 39374 GCAGGCACACCGCCGCCGC-ATCAGCGCGCAGGGCGTTCAGGTCGACGGG 2071ProLysHisLeuGluGluLeuAspLysSerHisLeuGluGlyGluLeuArgProLys	DD 39315 AACCAGAAGCGATCGCCGGAACGCCATCTCGAA

1487 ValhisProLeuAspValMetAlaAspAlaArgAlaLeuGluArc	40933 ACCAATCCACTTCCGACGTAGGTTCATCCACATGCAACGTCGCGGGCAGCTGCCCCG	GCATCGCCATCACCATCTTGATCACACCAGCGACACCCGCCGCCGCCT	1507SerLeuLysSerArgProGlyThrAlaSerSerSerGlyGlySerIleAlaAr;		AAGTGGCAATC	1546 H18G1yAlarrorneAlaG1yAlsheu	1562 MetArgGluProThrProArgLeuGlnGluGlySerLeuSerSerSerLysAlaSe	40663 CAGCACTCGCCAACGCCTGCGTGATCACCCGCTGCTGCGAGGGCCCA 1682 AsparoivsLeuthrSerThrProArgGlulleAlalysSerProHisSerThrVa		1602 GluHisHisProHisProlleSerProTyrGluHisLeuLeuAr			1638 ProhrgilyileProleuAspAlaAlaAlaAlaiyilyrivrieurioniyanshama 40487 CCCCAACCGGIGCCATCCGCAGCCGCCGCAAACGACTTGCACGGCCATCGGC		40430 AAGCCCCGTTGACGCGAAAACTCCACGAACATCCCCGGGCGTGGCCATCACCGTCAC	1667 ProfyrteulleArgdyJyfyrPloAspinfAladaueuGluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaueugluAsinfAladaue	1686 IleAsnAspTyrIleThrSerGlnGlnMetHisHisAsnThrAlaThrAlaMetAl	40310 ACCAACGACGA-CGAACACGCCGTATCCACCGTCACCGCAGGACCCTCGAAAA	40254 CGAATACGCAACCCGCAAAAACACGCCTCCCGCGCTGC-CATTV	1726 AlaGlyProArgGlyIleIleAspLeuSerGlnValProHisLeuProValLeuV		1/46 ProintPro	1749GlyThrProAlaThrAlaMetAspArgLeuAlaTyrLeuProThrAlaP:	40126 CGGTCCGCCCCCCCCGCAAAGCGGATCAATCCCCGCCCGC	1767 ProPheSerSerBerSerSerSerProLeuSerPro	40066 GACTTCCAGCAGCAACCGCTGCTGCGGATCCATCGCCAACGCCTCACGCGGGGAA
CGCGCCGGCGATGTC	othralaGinGlualaSerValLeuArgG 1197 	rileThrLysGlyIleProSerThrArgV 1217			/8GlyThrileThrArglielLeGlyGlud 1250 		NACT-CGCCGACCGATGGCCCAGCACCAC 41647	: cagcaaaccaacctgca	rGlyProProH	rccgarccaacaaggacccgrcggaaccaa 41537 spMetMetGluGlyargValGlyArgAlaI 1322	CCAGATGCGCGTCCAGTCCGGCACATGCTT 41477	1yargalaileFroProGlu 1338	roHisHisLeulysGluGlnH 1349	COACCCCCCCAGAGACCGGGATCCCGAC 41358			TCCGCTCCCA-GCACCGCCGCTCCTC 41266		sGlualaGlyArgSerlleHisGlulle	ACCCAAATAGGACAACAATGCGCTCGCCTG 41173		Tyrasp1460		ThrGlyAlaSerThrThrGlySerLysLysH1sAspval 14/3 		
41964 ATGCGAAACCGCAACGCCGCCACCGAATCCCTTGCCCATCCAGCGCGCGC	1177 lnAlaGlyProProGluSerLeuGlyValProThrAlaGlnGluA:	1197 lyThrAlaLeuGlySerValProGlyGlySerIleThrLysGlyIleProSerThrArgV	41856 G-ACCGCGGCGAIACCCGCACCAIC 1217 alProSerAspSerAlaileThrTyr	41806 GCTGCTCCACCAGCAGCCACCGCAACATGGCACCGCCAGAAGGCAACGCCTGCATCA	1230 hrHisGlyThrPrcAlaAspValLeuTyrLysGlyThTleThrArglleIleGlyGlUA 41746 ACCGGGCACGACCCCCACCATCCGAGCCGCATCGGGCAACG	1250 spSerProSerArgLeuAspArgGlyArgGluAspSerLeuProLysGlyHisVallleT	41704 ACAACAC-TCCGGCGCAACGGCGCGGGAACGGCGGAACGGAA	12/0 YICHUCIYABADSGIANISVALDGGGELYFGIGGTYFGI 11	1290SerLysGluAspGlyProH	41596 ACGCGAACAÀACCCGÀCTGGGCCCACAGCGTCCGATCCAACAAGGACCGGTGGGAACCAA 1302 isGluThrAlaAlaProLysArgThrTyrAspMetMetGluGlyArgValGlyArgAlal	:: ::	1322 leSerSerAlaSerIleGluGlyLeuMetGlyArgAlaIleProProGlu	1339		1349 isHisileArgGlySerlleThrGinGlyLleFroArgSerlyrvalGludiaGliblum 			1389 FOSETARGABBLEUITRELLATATYTUY INTELLATALEGUALYFLODGGAY GEGETATATYTUY INTELLATALEGUALYFLODGGAY GEGETATATYTUY INTELLATALEGUALYFLODGGAY GEGTATATYTUY INTELLATATATATATATATATATATATATATATATATATAT	1409 ro-AlaHisGluGlyLeuValAlaThrValLysGluAlaGly		1429 ProArgGluGluLeuArgHieThrProGluLeuProLeuAlaProArgFroLeuLyBellu 1172 AGCGCTGAGCGGTGCTCTTCCCCGACACGATCAACGGGACGGTGGAAGGTCTAC		41112 GGTCGATTCCGAATCGGGACCCTCTTGTCCGCAGTCTCGTCTGGTGTTTTCGAGGCTTG			1474 Argserbeulegiyserkrociyatginikme

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talagin 1705 ::: AACCAAA 40255 uValPro 1745 ||| ----CCT 40187 aProGln 1766 |||::: |CCCAAG 40067 uTyrPro 1666 : CACACCA 40371 nThrile 1685 ::: CAGCGCC 40311 ArgGly 1617 ||| |CGTTGC 40535 ArgAla 1502 ||||||| CGTGCC 40874 GluGlu 1506 :::::: GACCAA 40814 ArgGly 1525 ||| CGT--- 40757 TACCTA 40205 ACACGC 40127 HAAATCC 40007 ||| |GCCGCC 40431 CATTCG 40613 CCTTCT 40488 ProThr 1783 ACAAAC 40664 40592 TyrAla 1725 ----- 1748 CGGAT 40724 Serile 1637 AlaPro 1657 sluasp 1545 ValPro 1601 /alThr 1561 SerGln 1581

19954	Db 42043ValGlyProValThrMetGlyLeuProLeuProMetAspProLysL 1160 Qy 1145ValGlyProValThrMetGlyLeuProLeuProMetAspProLysL 1160 Db 42021 GCGTGCGATTTCGGCGAACTCCTGCAGTCCATCCGATACGAATGAAACGC 41965 Qy 1160 ysLeuAlaProPheSerGlyValLysGlnGluGlnLeuSerProArgGlyG 1177
633AssitypseralatiealaargmetvaldiysertysthrvalserdinCysiyahan 635 Priyrphehamytylysiyahangdinanieaasgdiutie	Oy 897 uGlyGlySerGLyArghlarhrAlaLysserSerSerGlyAlaProGlnAsp 914 bb 43074 ACCACGACGCTGAAGCCCCTGGAACACGCCCGTACTCGAACCGTTTCC 43015 Qy 915SerAspSerSerAlaThrCysSerAlaAspGluValAspGluAl 929 ti:

-60-SN	US-09-522-753-5 (1-2517) x AAF88313 (1-50000)	ò	324 AlaLysGluSerLysValArg
ò	ArgTyrProPro	व्य	::: 44994 ATAGTCGAACACCGCAGTCGC
Ор	45889 GAACCACCGCCAACACCCGATGCCCATTCCGCCGAGCATCCGACAACCGCTCCA 45836	δλ	343 GlnArgGluLeuGlnGluArg
ò	32 ArgThrHisThrAspValGlyLeuLeuGluTyrGlnHisHisSerArgAspTyrAla 50	qq	44934 CAACTCCACACCAGACAACGA
qq	45835 ACAACACCAAAC	ò	363 MetSerAlaAlaArgSerGlu
ò	51 SerHisLeuSerProGlySerIleIleGlnProGlnArgArgArgArgProSerLeuLeuSer 70	qü	
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ò	uGluProValSerProPro	q _Q	 44563 CCGTCATCCCGCTCGGCTGC
ф		ď	492 sSerGlnGlnGlnGlnGlnG
ò	152 SerProProHisThrAspProGluLeuGluLeuValProProArgLeuSerLysGlu 170	qq	44503 GTGACCGCCGGTAATGCGCC
Ωp		δ	512 tProArgSerSerGlnGluG
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qq	45375 CCGTTGCTGCGGATCCATCCC	ò	532 uGluGluLysProGluValG
ò	191 SerLysLeuLysLysElnGlnGlnLeuGluGluGluAlaAlaLysProProGluPro 210	QO	: 44383 TATCACGAGTCAGCTCGTGC
qq	45345 ACGCGGGGAAATCCCGAAGAACCCGCATCGAACCGATCGCCGTCGGTGAGGAAGGCCCC 45286	λ	552 spihršerGlyGluAspAsn
ģ	211 GluLysProValSerProProProlleGluSerLysHisArgSerLeuValGlnIle 229	đ	44323 CCACGTCGTCGCTGACAAC
qa		<i>ک</i> ر	sКП 295
ο,		අු	44263 CCACGCCGCACAACGGAAAT
qq	45255	ò	581 rglleThrArgSerMetAla
ò	250ProGlnValGluLeuProLeuTyrAsnGlnProSerAspThrArgGln 265	a a	
qq	45210 TGGGAAGTCCGCGACCGCATCCACGCCCCCCCCAAACGCCAGAGGTCCTCCGGCGA 45151	ò	593 luGluAlalleThrProGln
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à	304 GlnLeuMetGluAlaLeuGluLysLysValGluArgIleGluAsnAsnProArgArgArg 323	QQ —	:::
qa	45045 CTCGTCTGCCAGTTCCTGGTGCAGGAACCGGGCCAGCGCCTCGGCGTCGG 44995	_	

324	AlaLysGluSerLysValArgGluTyrTyrGluLysGlnPheProGluIleArgLys 342
е 4.	GlnargGluLeuGlnGluArgMetGlnSerArgValGlyGlnArgGlySerGlyLeuSer 362
363	MetSeralaalaargSerGluHisGluValSerGluIlelleAspGlyLeuSerGluGln 382 ::: grcaccccccccccccccccccccccccccccccccccc
383	GluabnLeuGluLysGlnMetArgGlnLeuAlaValIleProProMetLeuTyrAspAla 402
403	AspGlnGln-ArgIleLysPhelleAsnMetAsnGlyLeuMetAlaAspProMetLysVa 422 ::: ::: ::: :::
422	ITyrLysaspargGlnValMetAsnMetTrpSerGluGluGluLysGluThrPheArgGl 442
442	uLysPheMetGlnHisPro
452	YLEUIIEAlaSErPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyrTyrLe 472
472	uThrLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArgArgGlyLy 492
492	sserginginginginginginginginginginginginging
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552	spThrSerGlyGluAspAsnAspGluLysGluAlaValAlaSer566
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44083	CACCGCGTTCGGTCACCAGGTGGCGGGCAACCGCCCCCCCAACAGGCCCGGGGGGGG
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37789 GTAGCTGCCGCCTAAGCCGCCGCCGCCGTGCCG---GCGGGTGCCCCAGTGCGC 37845
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This invention describes a novel method nucleic acid (I) and its encoded polypeptide (II) containing at least one region that encodes an enzymatic activity involved in biosynthesis of spinosyns. (I) are used (i) to identify, inactivate or modulate genes involved in the biosynthesis of (II); (ii) to generate a library of polyketide synthases; (iii) for adding forosemaine or trimethylthamnose to a spinosyn or polyketide enzymes, which are used for production of (II), their precursors or derivatives, including production of (II), their precursors or derivatives, including production of (II), their precursors or derivatives, including production of (II), their precursors or and thus have increased resistance to insects. (I) are also useful as markers for sequencing of the Saccharopolyspora spinosa genome. (II) are markers for sequencing of the Saccharopolyspora spinosa genome. (II) are markers for used to raise specific antibodies, useful for identifying expression clones in a gene bank. Cells transformed with (I) may produce (II). This sequence represents a genomic DNA fragment of the S. spinosa genome which contains the coding regions for proteins involved in
                                    New nucleic acid encoding enzymes for spinosyn biosynthesis, useful for recombinant production of insecticidal spinosyns and their derivatives.
2486 aGlySerGlyProLeuAlaGlyProHisHisAlaTrpAspGluGluProLysProLeuLe
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Forosamine, trimethylrhamnose, polyketide synthase, biosynthesis, spinosyn; polyketide aglycone, transgenic plant, insect resistance; macrolide, insecticidal, ds.
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1480 oGlyArgThrPheProProValHisProLeuAspValMetAlaAspAlaArgAlaLe 1499	CGGGGGCGCGTTGGCGGTTGCCGATCAACGGGCGCCCGGTATTCGCAGAAGAACT uThrTyrGluAspHisGlyAla		1589 OLEUAIDE AND SOLITION SECURIFICATION SECURIFIC	1649 TTYLLGUPLOARGHisLeuAlaProASDPTOThrTYPPTOHI 1663 36359CTGCCGCGATAACCTCTGGCGCCGCAATCACAACGACATCTGACACTCTGACACTCCCAATA 36414 1663 SLeuTyrProProTyrLeuIleArgGlyTyrProAspThrAlaAlaLeuGluASDATG - 1682 1663 SCATGACGGCTCTGTCATGCGGCCCGGGAACGTCACCACCAGTAAAATCGGCA 3648 36415 GCATGACGGCTCTGTCATGCGGACCGGGAACGTCACCACAGAAAATCGGCA 36468 1683	1687 nAspTyrileThrSerGlnGlnMetHisHisAsmThrAlaThrAlaMetAlaGlnArgAl 1707 : ::::: 36529 cGACTTCGTGTACCGGGTGAACCCGGTGTGTTTTGAAGGACAGCCCGC 36576 1707 aAspMetLeuargGlyLeuSerProArgGluSerSerLeualaLeuAsmTyrAlaAlaGl 1727 :::::
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36779	::: cgccattgccgccggtaccggcgccaaac 368
1805	pArgAspArgGluArgGluLysSerIleLeuThrSerThrThrThrValGluHis-AlaP 1825
1825	rolleTrpArgProGlyThrGluGlnSerSerGlySerSerGlySerSerGlyGlyGlyGl945
1845	1yGlySerSerSerArgProAlaSerHisSerHisAlaHisGlnHisSerProIleSerP 1865
1865	roArgThrGlnAspAlaLeuGlnGlnArgProSerValLeuHisAsnThrGlyM 1883
1883	etLysGlyIleIleThrAlaValGluProSerLysProThrValLeuArgSerThrSerT 1903 ::: GTAGGCGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGG
1903	hrSerSerProValArgProAlaAlaThrPheProProAlaThrHisCy8ProLeuGlyG 1923 :::
1923	lyThrLeuAspGlyValTyrProThrLeuMetGluProValLeuLeuProLysGluAlaP 1943
1943	roargValAlaangProGluArgProArgAlaAspThrGlyHisAlaPheLeuAlaLysP 1963
1963	roproAlaargserGlyLeuGluProAlaserSerBroserLysGlySer 1979 ::: ::: cGccGCCGCCGCCGCCGTTGGCGAACAGCCCACCGG 37275
1980	GlubroArgProLeuValProProValSerGlyHisAlaThrIle
1995	AlaargThrProAlaLysAsnLeuAlaProHisHisAlaSerProAspP 2011
2011	roprohlaproprohlaserhlaserhsprohlsargglulysThrGlnserLysProP 2031
2031	heSerIleGlnGluLeuGluLeuArgSerLeuGlyTyrHisGlySerSerTyrSerProG 2051 :: TCAGCAAGGCGGCTC 37458
2051	luglyValGlubroValSerProValSerSerProSerLeuThrHisAspLySGlyLeuP 2071
2071	rolysHisLeuGluGluLeuAspLysSerHisLeuGluGlyGluLeuArgProLysGlnP 2091
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114	ovalThrMetGlyLeuProLeuProMetAspProLysLysLeuAlaProPheSerGlyVa
35127	
1167	ILysGlnGluGlnLeuSerProArgGlyGlnAlaGlyProProGluSe 1183
118	rleuGlyValProThrAlaGlnGluAlaSerValLeuArdGlyThrAlaLeuGlySerVa 1203
35243	
1203	IProGlyGlyGlyCleThrLysGlyIleProSerThrArgValProSerAspSerAlail 1223
35303	
1223	eThrTyrArgGlySerlleThrHisGlyThrProAlaAspValLeuTyrLysGlyThrIl 1243
35360	
1243	eThrarglleIleGlyGluAspSerProSerArgLeuAs
35376	GAGGCGAGCATGCCGCCAT 35395
1263	uProLysGlyHisVallleTyrGluGlyLysLysGlyHisVa
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1283	yMetSerValThrGlnCysSerLysGluAspGlyArgSerSerSerGlyProProHi
35444	TGCCGCCTTCCCCGCCTGCCCGAACA 35470
1303	uThralaalaaroLysArgThrTyrAspMetMetGluGlyArgValGlyArgAlaileSe 1323
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1343	sHisLeuLysGluGlnHisHisIleArgGlySe
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1400	nAlaLeuGlyProLeuLysLeuLysProAlaHisGluGlyLeuValAlaThrValLysGl 1420
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g (10.00 0.0000000000000000000000000000000	Qy 747 pThrGluSerIle
& 1	I.eAlaSerPheLeuGluArgLy8ThrValAla	DP 33909 GCCGCCGTTCCCG
g (32952 GICGCAGAGIIIICGCAGCAGCAGCAGGGGGCCGGGGGCLIGICGGAAAGGGGGGGGGG	Qy 767 oLysProProAla'
Š		Db 33933 dececcerinec
3 8	LysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArg4	
. q	33072 GAAGATGCGTCGTTGTTAGCCGGAGGAGGACGCCCACACTAGGTTCGGCACTGT 33131	33993
ò	491GlyLysSerGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnG	gg A
qq	33132 GAAGGGGTCGTCGGCCGCAAGCAGTCGATCGAACCAGGGGCGGACGGTTCGCTGATTCGG 33191	
ò	510 GlnProMetProArgSerSerGlnGluGluLysAspGluLysGluLysGluLys 527	

33192	ATGGTCACCGCGGTGCAGCCAGGAGGTCGATGAGGAACATCGCCTATTT 33251
528	
33252	GIGCCIGICCAGGCICACTICCGCGAGIICAGIICCAGACCCICGICGAGCACIICGGAC 33311
545	LeuLeuLysGluLysThrAspAspThrSerGlyGluAspAsnAspGluLysGluAlaVal 564
33312	AACACCGTATTCGAGGTTAGGTCGATACCTGGCCGCGGACCGGTGCCGGCGTCAAAAACG 33371
565	AlaserLysGlyArgLysThrAlaAsnSerGlnGlyArgArgLysGly 580 :::
581	ArglleThrArgSerWetalaAsnGluAla 590
33432	
591	ST.
33492	
33552	
607	tGluLeuAsnGluSerSerArgTrpThrGluGluGluMetGluThrAlaLysLysGlyLe 627
33612	::: GCGCAAGCGTTGAGTCCGTTCTCGCCGAGCAGCCGCCGG 33654
627	uLeuGluHisGlyArgAsnTrpSerAlaIleAlaArgMetValGlySerLysThrValSe 647
33655	-IGCCGCCGGCACCGGGCGTGGCCCCCGGCTTTGCCGG33690
647	rGlnCysLysAsnPheTyrPheAsnTyrLysLysArgGlnAsnLeuAspGlulleLeuGl 667 ::: ::: -:GGTTGCCGCCGTTGCCGCGTTGCCGATCAGCACGGCGTTGCCGCCG 33737
199	nGlnHisLysLeuLysMetGluLysGluArgAsnAlaArgLysLysLysLysAlaPr 687
33738	
687	OAlaAlaAlaSerGluGluAlaAlaPheProProValValGluAspGluGluMetGluAl 707
33795	gTTGCCGAACACCCCGGCGTGGCCACGGTTGCCGAACACCCCGGCGACGG
70	SerGlyValSerGlyAsnGluGluGluMetValGluGluAlaGluAlaLeuHisAlaSe
33823	3386
	rGlyAsnGluValProArgGlyGluCysSerGlyProAlaThrValAsnSerSerAs 747
9	GGCACCGCTGCCGCCGCCGGCGCCGCCGCCGCCGGCGCGAAGCCGAAGCAA 33
74	pThrGluSerIleProSerProHisThrGl
33909	GCCGCCGTTCCCGCCGGCCCG
33933	oLysproProAlaThrLeuGlyAlaAspGlyProProProGlyProProTh 784
784	
33993	::: GCCGCCGAAGCCGAAGAGTAAGGCGCCGTTCCGCCGGCCCGGCCGG
0	y81aProThrProProAlaProProSer
34053	GGAGCTGGCGCCACCGCTGCCGCCGCCCACCGGAGGCGCCCGAGGGAGAGTAGGCCGCGG
815	ProSerAlaProProProValValProLysGluGluLysGluGluThrAl 832

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Db 41746 CCGCCGTTACCGCCGGCACCGTTGTTGACGGCGAAACTCGGATCGCCGCCGCCCCTTA 41805 Qy 2256ArgMetGlySerLysSerProGlyAsnThrSerGlnProProA 2270	2310 lyThrGluIlePheAsnMetProAlaIleThrGlyThrGlyLeuMetThrTyrArgSerG 233: 41968 GCGCCGCCGTCCCGAACAGCCCGGCGTCGCCACCGTTGCCGCTTACC	Qy 2406 erGlyArgPrSerSerArgLysAlaLysSerProAlaProGlyLeuAlaSerGlyAspA 2426 bb 42190 ccGTTCACCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCC	2505 euLeuCysSerGlnTyrGlu 2511

8 8	3/888 GCCACCGIGGCCGCGGAGCCGGCCGCCGCGGCGGCGGCCGCCGGGGGG	ò	1340 isSerProHisHis
qq	37946 AGGCCACCGTGGTCACCGTTGCCCCCGTCGCCGCCTTGCCGCCAGCAGCACGCCGGT 38005	qo ,	
ò	1027 aProProAlaAspLysGluAlaPheAlaAlaGluAlaGlnLysLeuProGlyAspProPr 1047	हें व	1360 roArgSerTyrVal
a	38006 GCCGCCGGCGCCGCCGTTGCCGCCGTTGCCGCCGCGCGCG	8 8	38912 CGAAGGAC=====
È	OCYSTrpThrSerGlyLeuProPheProValProProArgGluValIleLysAlaSerPr	3 8	38921 CGCCGGCGACACCC
d d	38048 GCCAATGCCGCTGCCGCTGCCCCCGGTGCCACCGAACCCACCTGGCCCACGTGCGCC 38104	ò	1398 ysThrGlnAlaLeu
à i	OHisAlaProAspProSerAlaPheSerTyrAlaProProGlyHisProLeuProLeuGl	i 8	
g (GCCGGCCGCCGCTGCCGCTGCCGCCGCCGCCGCCGCGCGCCGC	ò	1418 alLysGluAlaGly
è	108/ YLEUMIBABDINIALARGYROVALLEU	qq	39005 CGGAGCCGGCG-
8	-ProArgProProThr1]eSerAsnProProProLeu1]eSerSerAlaLv	ò	1438 luLeuProLeuAla
; a		qq	39047 cerracceccade
è	BHisProSerValLeuGluArgGlnIleGlyAlaIleSerGlnGlyMetSerValGlnLe	λõ	
q	38282 TGAGCCGGCGTTGCCGGCATCGCCACCGCTGCCGCC 38323	අු	39095 CGATACCGCCATTC
Ś	uHisValProTyrSerGluHisAlaLysAlaProValGlyProValThrMetGlyLeuPr	δ	1478 GlySerProGlyAz
: A		q	ငဇငင်
ò	OLEUProMetAspProLysLysLeuAlaProPheSerGlyValLysGlnGluGlnLe	ò	1491AspVe
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ે ઠે	uSerProAradlyGlnAlaGlyProProGluSerLe	ò	1508 LeuLysSerArgP1
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Š	_	č	1527 ovalilevalProC
· 원		අු	39312 AAACCCAGTCTCGC
8	_	δλ	1546 BGlyAlaProPhe/
; A		Q	39372 GCCGCCACCGC
l 8	_	δ	1566 rProArgLeuGlnC
S 8	•	අු	39432 GCCCCTGGCCG
à	sGlyThr1leThrArg11e11eGlyGluAspSerProSerArgLeuAspArgGlyArgGl	ò	1586 rSerThrProArg(:::
셤	38667GCCGTGGCCGCTGCGGCGCTGCCGCCGCCCCCTGGCCGCC	q 0	
ò		Š	1606 sProlleSerProl
අු		q	39552 GCCACCCAACCCG
ò	1280 rGluGlyGlyMetSerValThrGlnCysSerLysGluAspGlyArgSerSerSerGlyPr 1300	Š	rHisIl
g	38768 GGCCGGCGCTGACGTTGACGACGCCTGAGCCGCGGGAC 38809	q G	
ò	1300 oProHisGluThrAlaAlaProLysArgThrTyrAspMetMetGluGlyArgValGlyAr 1320	87∵ 1	1645 aAlaAlaAlaTyr7
đ	38810 CGCTGCTGCCGTTGCCGCCTTTGCCGCGCGCGC	g è	19618 GGCGACGTCAAG
ઠે	leGluGlyLeuMetGlyA	ÿ £	2000 4000000000000000000000000000000000
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1340	isSerProHisHisLeuLysGluGlnHisHisIleArgGlySerIleThrGlnGlyIleP 1360
1360	roArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGluAlaLysLeuLeuLysA 1380 ::: CGAAGGACA 38920
1380	ProProProProPro GGGTTGCCGCCGGCCA
1398 38959	ysThrGlnAlaLeuGlyProLeuLysLeuLysProAlaHisGluGlyLeuValAlaThrV 1418
1418	ysgluðlag :: AGCCGGCG-
1438	InleuProLeuAlaPro-ArgProLeuLysGluGlySerIleThrGlnGlyThrProLeu 1457
1458 39095	LysTyrAspThrGlyAlaSerThrThrGlySerLysLysHisAspValArgSerLeulle 1477 ::: CGATACCGCCATTGCCGCCCGCCGCGGAGCCGGCGTCAC 39136
1478	52555
1491	
1508	LeuLysSerArgProGlyThrAlaSerSerGlyGlySerIleAlaArgGlyAla-Pr 1527
1527	oValileValProGluLeuGlyLysProArgGlnSerProLeuThrTyrGluAspHi 1546
1546 39372	### ##################################
1566 39432	rProArgLeuGlnGluGlySerLeuSerSerSerLysAlaSerGlnAspArgLysLeuTh 1586 GCGCCCTGGCCGCCGACCCCCGCCGGCACCGCCGCCGCCGCCGCCGC
1586 39492	rSerThrProArgGluIleAlaLysSerProHisSerThrValProGluHisHisProHi 1606 :::
1606 39552	sProlleSerProTyrGluHisLeuLeuArgGlyValSerGlyValAspLeuTyrArgSe 1626
1626 39568	roThr cGGCACCG
1645 39618	aAlaAlaAlaTyrTyrLeuProArgHisLeuAlaProAsnProThrTyrProHisLeu 1664 ::: GGCGACGTCAACGGCCGCCACCGCCGCCGAAGCCGCCGAGGCCGCGTTGAT 39677
1665 39678	

AAI99682_22	1988 CAGÁTCTCCAGCAGCCGGAACAGCGCCACCTCGAAGGCGAACAGGCGGGCTGGGCGAAA 1929	-
AAI99682 19 AAI99682 20 AAI99682 21	/ 2282 MetValLysSerLysLysGlnGlulleAsnLysLysLeuAsnThrHisAsnArgAsnGlu 2301	•
AA199682 1/ AA199682 18 AA199682 19	2034CAGTTCACCGACGGAATGGCCGAAGGACGTAGTCGGGCGTCAGACCC 1989	-
AA199682 15 AA199682 16 AA199682 17	/ 2262 roGlyAsnThrSerGln-ProProAlaPhePheSerLysLeuThrGluSerAsnSerAla 2281	-
AA199682 13 AA199682 14 AA199682 14	2090 ACCACCAGGACGCACGCAGGAAGCAGCGACATCCCGGCGACATGGGCGGCGGCGGCGGC 2035	-
WP AAI99682_11 110 WP AAI99682_12 120 WP AAI9682_13 130	2242 yrProLeuLeuTyrArgAspGlyGluGlnThrGluProSe	J
AAI99682 09 AAI99682 10	2143 TGGCCACCATGGTTCCGCCGGACGGGAGCCTTG-CATCAGGCGCCTC	-
AA199682_07 AA199682_08	2222 leGluProValSerProProGluGlyMetThrGluProGl	<u>.</u>
AA199682 AA199682	2176 GCAGCGGGAGG	-
AA199682_03 AA199682_04	2202 lyGlyLysArgSerProGluProAsnLysThrSerValLe	•
AAI99682_02	22316 GGACGATCGACCGCGGCCCGTTCACCGCGGCGCGCGCGCG	-
AA199682 00	/ 2186ProProAspHis-GlyAlaProAlaArgGlySerProHisSerGluG 2202	~
Sequence split int	2296 decedentecececentracracecaagarecaadacecarecreacaaa 2237	-
37 ion (38 of 45) of	/ 2169 aSerCysProValLeuAspLeuArgArgProProSerAspLeuTyrLeu 2185	Ŭ
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1460 CACGGC	Db 2377 CGCCGCGCGCGCGCAAGTCGCCCAACATGGCGTCCATGTG	
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Qy 2435 rGluGlyAspCysA	2112 rG]nbroSerSerSerBro[.e.](anifaniG]nThr&]aProG]vWallvsG]vHi 212	
٠ ن	2557 CGATTTCGAGGCAGGTGACGCCCTGTTCCTGAAGCCATGAGATGCCGTCCGCGAAGAAGC 249	, u
Qy 2415 sSerProAlaProG		- '
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2376		J
1754 †ĠĠGTTCC	2730 AGCAGTCCCTCGAACACAGGGCCACTCAACCGCCGTCCCAGC-CACGAACA 267	- 14
Qy 2356 GlnTrpGluGluSe		_
1808	2783CGCTCAAACGCATACGTCGGCAGCT	1
2336	2036 uGluLeuArgSerLeuGlyTyTHisGlySerSerTyrSerProGluGlyValGluPr 20	
Db 1868 AGCCAGGGGCACC	2808 CTCCACACACACACACACACACACACACACACACACACA	, L
Qy 2316 MetProAlaileTh	2000 CGC-LICCALANGGSACCACCACCACGAAAACCCCCGGAAACCICCGGAAACCICCGCAGAILCCCIC 200	. (
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Qy 2302 ProGluTyrAsnIl		

AA199682 from base 3700001 (Mycobacterium tuberculosis strain ragments LOCUS AA199682 Accession Aa199682 GCCCGGCGAAGAGT---ACGGCGAGTTCGCCC-----GAGGTG 1710 AsnArgArgThrProLeuThrAsnArgValTrpGluAspArgProSe 2455 ||||||:::||| TCAGCCGGCCTTCGGGAGAGTCCTGCGGTGCGAAGAGTACGTCCCGC 1869 erProProLeuSerAlaAsnAlaPheAsnProLeuAsnAlaSerAla 2375 laMetProlleThrAlaAlaAspGlyArgSerAspHisThrLeuThr 2395 SGlyLeuAlaSerGlyAspArgProProSerValSerSerValHisSe 2435 GCCCGACCCCAGCGGACTCACCCCACATCCGCACCCCCAC-- 1532 ThrproPheProTyrAsnProLeuIleMetArgLeuGlnAlaGlyVa 2475 ProProProGlyLeuProAlaGlySerGlyProLeuAlaGlyProHi 2495 ---IlePheAsn 2315 hrGlyThrGlyLeuMetThrTyrArgSerGlnAlaValGlnGluHis 2335 cedestresersesses estates contracted contracted the contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contract contr letGlyLeuGluAlaIleIleArgLy8AlaLeuMetGlyLy8TyrAsp 2355 llyGlyLysAlaLysValSerGlyArgProSerSerArg-LysAlaLy 2415 cegaaceedacaccaeecegacccaacteergeageegeegeega 1401 AspGluGluProLysPro 2504 acccccaccccrcca 1370 leserGlnProGlyThrGlu--110000 310000 410000 510000 710000 1010000 11110000 11210000 11410000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 11710000 fragments Begin 1000001 2000001 2000001 2000001 2000001 2000001 2000001 2000001 2000001 2000001 2000001 2000001 2000001 2000001 2100001 2200001 2300001

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ACTICACGCICGIGCCGIACCCCCATICCCCCAACGCCGACACCACCACCCAGC 2928 pGlyValTyrproThrLeuMetGluProValLeuLeuProLysGluAlaProArgValAl 1946 ------chactccacccaakca---chacaccccacaccdchcaacccctc 3108 |||| |:::|||| GCCACCACTCGCCTCGGACACCACCACGCGCGCCCGACAGC 3048 oAlaArgSerGlyLeuGluProAlaSerSerProSerLysGlySerGluProArgProLe 1984 ------LysProThrValLeuArgSerThrSerThrSerSerProValArgPro 1909 AlaAlaThrPhePro------proAlaThrHisCysProLeuGlyGlyThrLeuAs 1926 3420 SerArgProAlaSerHisSer----HisAlaHisGlnHisSerProIleSerProArg 1866 43 LeuValProProThrProGlyThrPro---AlaThrAlaMetAspArgLeu---AlaTyr 1760 76 AAGTCCGCTCGGAGTTCTTCGACTCCTCGGGGGCTCTCCCCGCGTCGGCTCACCAGC 3717 61 LeuProThrAlaProGlnProPheSerSerArgHisSerSerSerProLeuSerProGly 1780 :::||||||||||||1|||16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 16 || 81 GlyproThrHisLeuThrLysPro------ThrThrThrSerSerSer 1794 95 GluargGluargAspargAspargGluargAspargAspargGluargGluLysSerIle 1814 LeuThrSerThrThrThrValGluHisAlaProlleTrpArgProGlyThrGluGlnSer 1834 ThrGlnAspAlaLeuGlnGlnArgProSerValLeuHisAsnThrGlyMetLysGlyIle 1886 ---Aceccacatracchaccactecaccacactacaccacacacteccactcccactecac 30 ICACGGTCGCCGCTCACCGCCACCGAC---ACCCGCGCGCCCCAAG---CCCATC 3777 -----ArgAlaAspThrGlyHisAlaPheLeuAlaLysProPr CCCCCTACACCACCACGACGACGACGACTCCCGATAGCGCCACCGGTCGACGC uValproprovalserGlyHisAlaThrIleAlaArgThrProAlaLys------260 ACGCCACTCCCCCAGACGCAGAGGAGGACGACCGCCGACGACAGCTCACCATCGACGCC 23 AsnTyrAlaAlaGlyProArgGlyIleIleAspLeuSerGlnValProHisLeuProVal 79 -----CAGCGGCA-----CTCCGCCGGAACGCCGACGCAACACCTCCGCCAACGCC 29 CCCTCCAACACCCCGTCATCCACAACTCCCGCGGGATGCACCACCACCACCACC------03 etalaGlnArgAlaAspMetLeuArg-GlyLeuSerProArgGluSerSerLeuAlaLeu 946 aArgProGluArgPro---------387 IleThrAlaValGluProSer------910 047 154 952 335 198 359 894

1046 ProProCysTrpThrSerGlyLeuProPhèProValProProArgGluValIleLysAla 1065 	1066 SerproHisAlaProAspProSerAlaPheSerTyrAlaProProGlyHisProLeuPro 1085	1086 LeuGlyLeuHisAspThrAlaArgProValLeuProArgProProThrIleSerAsnPro 1105	1106 ProProLeulleSerSerAlaLysHisProSerValLeuGluArgGlnIleGlyAlaile 1125 5551 CCGCCAAAA 5543	1126 SerGlnGlyMetSerValGln-LeuHisValProTyrSerGluHisAlaLysAlaProVa 1145	1145 1GlyProValThrMetGlyLeuProLeuProMetAspProLysLysLeuAlaProPheSe 1165	1165 rGlyValLysGlnGluGlnLeuSerProArg	1176GlyGlnAlaGlyProProGluSerLeuGlyValProTh 1188	1188 ralaGlnGlualaSerValLeuArgGlyThr-AlaLeuGlySerValProGlyGlySerI :::	1208 leThrLysGlylleProSerThrArgValProSerAspSerAlaIleThrTyrArgGlyS :::	1228	1242 hrIleThrArglleIleGlyGluAspSerProSerArgLeuAspArgGlyA 1259	1259 rgGluAspSerLeuProLysGlyHisValIleTyrGluGlyLysLys- 1274	1275 GlyHisValLeuSerTyrGluGlyGlyMetSerValThrGlnCysSerLysGluAspGly 1294	1295 ArgSerSerSerGlyProProHisGluThrAlaAlaProLysArgThrTyrAspMetMet 1314	1315 GluGlyArgValGlyArgAlalleSerSerAlaSerIleGluGlyLeuMetGlyArgAla	4941 1335	4940GAACCCCCGCCCCGCTCCGGATCATACAAAGCCTCCACGTCCCAACCCCGGT 4886 1354 rlleThrGlnGly1leProArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGl 1374	4886	1374 uAlaLysLeuLeuLysArgGluGlyThrProProProProProSerArgAspLeuTh 1394
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ACCCATCTCGTCCAGGGTTTCCTGCACCCGCGACCGCCGG 7225 |||:::||| -----CACGCCTCGATGAAGACGGT 7198 ::: -----GICGCGICCCGGAAGCGGACCTGGCGCAGGIIGCG 7135 |||::: |GGTGGCAAGCGGCCTTCGCCC---CGGCGCAGAGATCCGACGAC 7285 rGluserIleProserProHisThrGluAlaAlaLysAspThrGl 763 -----ProAlaThrLeuGlyAlaAspGlyProProProGlyProP 783 Argibiser---- 790 ::: segradcargecegegeacgergrecacegecrecegacaacre 7522 .:: ||| |||||| ::| NGAGACCCCGGAACCCCGCGCGCGCCCTCCACATCCAGCCAAAA 7462 odcaracercescente de 1402 de 1402 de 1402 de 1402 de 1402 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1403 de 1 uGluHisGlyArgAsnTrpSerAjaIleAlaArgMetValGlySe 643 nHisLysLeuLysMetGluLysGluArgAsnAlaArgArgLysLy 683 aAlaAlaSerGluGluAlaAlaPheProprovalValGluAspGl 703 rgjyvalSerGlyAsnGluGluGjuMetValGluGluAjaGluAl 723 yAsnGluValproArgGlyGluCysSerGlyProAlaThrValAs 743 uLeuAsnGluSerSerArgTrpThrGluGluGluMetGluThrAl 623 nÇysLysAsnPheTyrPheAsnTyrLysLysArgGlnAsnLeuAs 663 AlagluLysglugluLysProgluValGluAsnAspLysGl 543 ılysThrAspAspThrSerGlyGluAspAsnAspGluLysGluAl 563 nGluAlaAsnSerGluGluAlaIleThrProGlnGlnSerAlaGl 603 ndinginprometProArgSerSerGinGluGluLysAspGluLy 523 yargıysThralaAsnSerGlnGlyarqArqArgLysGlyArgIleTh ıTyrLysSerLeuValArgArgSerTyrArgArgArgG|YLysŞe GlnGlnGlnGlnGlnGlnGlnGlnGln-------AGGTCCAGCACGGTCTCGGCCTGTGCG-

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Alignment Scores:
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                                                                                                       GCCTGGGCGAAGACGGGGTACGCCTCGTACAGTTCGCGGCCCATGCCTGCGCGC----- 1755
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                                                                           Avermectin aglycone synthase; AAS; avermectin derivative;
drug production; vetinary drug; pesticide; ds.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                       The present invention relates to the production of modified derivatives of avermectin aglycone synthase (AAS) derived from Streptomyces evermitilis. The activity of an acyl carrier protein (ACP), beta-ketoacyl carrier protein synthase (KS), acyltransferase (AT), beta-ketoacyl carrier protein reductase (KR), dehydratase (DH), enoyl reductase (ER) and/or thioseterase (TB) domain may be reduced or suppressed. The process can be used in the production of drugs, vetinary drugs and pseticides. The present sequence is a fragment of the S. avermitilis genome
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8674 GATGACCACGACGTTCAGCCTCGGACAGCCGCTCCACCACCACACCACCACCACCTCGG
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                                                                                                                                                                                                                                                                                                           modified avermectin aglycone synthase derived from Streptomyces rmectilis used in production of 22,23-dihydroavermectin Bla used
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Mismatches:
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1684 hrileileasnaspTyrileThrSerGlnGlnMetHisHisAsnThrAlaThrAlaM 1703 3929 CCCTCCAACACCCGTCATCCACACGCGGGGTATGCACCACACACCACACCCACACACCACACACA	S a S
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1743 LeuValProProThrProGlyThrProAlaThrAlaMetAspArgLeuAlaTyr 1760 	3 A &
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1781 GlyproThrHisLeuThrLysProThrThrThrThrSerSer 1794	2 A S
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2090 ACCACCAGGACCACCGCGAGGCAGCAGCATCCCGGCGACATGGGCGGCCGC--- 2035 2177 2144 2222 leGluProValSerProProGluGlyMetThrGluProGlyHisSerArgSerAlaValT 2242 yrProLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlySerLysSerP 2262 2282 MetValLysSerLysLysGlnGluIleAsnLysLysLeuAsnThrHisAsnArgAsnGlu 2301 2129 2378 ------ProProAspHis-GlyAlaProAlaArgGlySerProHisSerGluG 2202 2202 iyGlyLysArgSerProGluProAsnLysThrSerValLeuGlyGlyGlyGluAspGlyI 2222 2130 -GlnArgValValThrLeuAlaGlnHisIleSerGluValIleThrGlnAspTyrThrAr 2149 2296 decegectrosteccoccatitlecroclobanderccadalcocostecticorccoognan 2237 2075 2618 2092 2112 rGlnProSerSer-----SerProLeuGeuGlnThrAlaProGlyValLysGlyHi 2129 2016 2809 2016 aserAlaserAspProHisArgGluLysThrGlnSerLysProPheSerIleGlnGluLe 2036 2055 ------CGCTCBACGCATACGTCGGCACCCCCGCATCCATCACCGACCTCGCG 2731 2730 AGCAGTCCCTCGAACACACGGCC----CACTCAACGGCGTCCCAGC-CACGAACA 2678 2091 2617 TESCECEGECCEGIECACTEAGCEAGTCCTGTECGAGGECCGACAGGCGTGCCGTCGGGGC 2558 2262 roGlyAsnThrSerGln-ProProAlaPhePheSerLysLeuThrGluSerAsnSerAla 2236 ggacgarcgaccdccggcccgrrrcaccgccgcdcgcgcccgggargcgrarrcgrcg 2075 uGluLeuAspLysSerHisLeuGluGlyGluLeuArgProLysGlnPro-----2092 yProValLysLeuGlyGlyGluAlaAlaHisLeuProHisLeuArgProLeuProGluSe 2437 GGGGCGCCCGGTGACGTTGGAGGACGACGGGAATCCGCGGGGGCGCTGAAGGTGACCTGCT gHisHisProGlnGlnLeuSerAlaProLeuProAlaProLeuTyrSerPheProGlyAl aSerCysProValLeuAspLeuArgArgProProSerAspLeuTyrLeu------2055 oValSerProValSerSerProSerLeuThrHisAspLysGlyLeuProLysHisLeuGl 1984 uValProproValSerGlyHisAlaThrIleAlaArgThrProAlaLys-----------AsnLeuAlaProHis---HisAlaSerProAspProProAlaProProAl 2036 uGluLeuhrgserLeuGlyTyrHisGlySerSerTyr---SerProGluGlyValGluPr 2129 B-----2186 2176 2169 2557 2783 2092 2927

ualalysLeuLeuLysArgGluGlyThrProProProProProSerArgAspLeuTh ::	ulysGluGlyserlleThrGl GCGGAGGAGTGGGGTAGTCGAAGATCAGCGTGGTGGCGGGAGGAGCCCGGTCGTCG GCGGAGGAGTGGGGTAGTCGAAGATCAGCGTGGTGGCGGCAGGAGGAGCCCGGTCGCGG GTTGAGGCGGTTGCGCAGTTCGACGGCGCTCACGGAGGAGCCCAGGTCGCGGAAGGC IATGAGGCGGTTGCGCAGTTCGACGGCGCTCACGGAGCCCAGGTCGCGGAAGGC IATGAGGCGGTTGCGCAGTTCGACGGCGCTCACGGAGCCCAGGTCGCGGAAGGC IATGAGAAGAGCGCTCACGGAGGCGGTGGCTGCTCCCGAGCACGGT IMELALAASPALAATGAAAACGGCGGTGGGGGGTGCTCTCCCGAGCACGGT IMELALAASPALAATGAAAACGGCGGTGGGGGGTCCTCCGGGGGGGTGCTCTCCGGGCGGGTGCTCTCCGGGCGGGGGG	1509 8SerMagProGlyThrAlaSerSerGryGlySerIleAlaArgGlyAAaProva 1528 1518	### SECTOR TO A SECTION OF THE FORM OF THE FOLLOWING SECTION OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF THE FORM OF TH
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đ	7635 GGCATGGGTAAAGGACCAGGTCCAGGACTCTGGCG 7593	È	791ArgAlaProlle
ò	473 rLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArgArgGlyLysSe 493	S 8	GAGGGAGAGTGCACCGGCG
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වූ	ACCAACACTCCCGAACACCGGAACCACCCACCCACCCACC	ð	871 lu
λ	563 aValAlaSerLySGLyArgLySInTAlaAshSeCINGLyArgArgLySGLYArgLielli 303	Ωp	6439 AGTGGGTGCCTGTCCTCCGAACAGCATGGCC
අ	CACGGACATCAGGAAGCGTCGCAAGCCGCCTTCGCCCCGGCGCAGAGATCCGAACAAC	δ	876 ysGlyLysAspAlaGluAlaAlaGluAlaTh1
ò	rargSerMetAlaAsnGluAlaAsnSerGluGluAlalleinrFrOGluGluGluSrAsdra 	qq	6379 CACCCTGTACCACCCCCGCGGGTGGCTCCCC
đ	CAGGCTGTCCAAGTCACCCATCTCGTCCAGGGTTTCCTGCACACCGACCG	λΌ	896 ysGluGlyGlySerGlyArgAlaThrThrAla
ò	erMetGluLeuAsnGluSerSerArgTrpThrGluGluGluHetGluThrAl	qa	6319 ACTCGTCGCGGTCCTCGGCCACGACCACGC
අු	CACGCCTCGATGAAGACGGT	ò	916 spserserAla
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ò	643 rLysThrValSerGlnCysLysAsnPheTyrPheAsnTyrLysLysArgGlnAsnLeuAs 663	g	6202 CCGCCACACCCCACGCAACCGACGCGCCTG
οg	7187CGGACCAGCGCCTGT7173	ò	927 spGluAlaGluGly-GlyAspLysAsnArgL
ò	663 pGlulleLeuGlnGlnHisLysLeuLysMetGluLysGluArgAsnAlaArgArgLysLy 683	셤	
qq	7172	ò	947 proThrGlyAspproArgAlaAsnAlaSer
ò	SLysLysAlaProAlaAlaAlaSerGluGluAlaAlaPheProProValValGluAspGl	qg	6091 CCTGCAGCCGGCCCGCACCCCCATCCGCGCC
අු	7134 GTACCAGTAGTCGGCGTCGAGTTCGGTGCCGTCGATCGCTTCGCCGGTGACGGTGAGTA 7075	ζ	961LeuAspLeuLysGlnLeuL
ò	703 uGluMetGluAlaSerGlyValSerGlyAsnGluGluGluGluMetValGluGluAlaGluAl 723	q	6031 CATTCGTCCCACTCACCCCGAACGCAGACAC
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ò	aLeuHisAlaSerGlyAsnGluValProArgGlyGluCysSerGlyProAlaThrValAs	qa	5971 CCGGCCACTCCACCTCATCCGCCAACACACG
අු		ογ	266
ò	743 nAsnSerSerAspThrGluSerIleProSerProHisThrGluAlaAlaLysAspThrGl 763	qa	5911 ACGACGGCTCATCCACATGCAACGTCCGCGG
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ð í	770	qq	5791 ACCCCAACCACACGGCCGCTCCCCGAACG
Q C	6849 CCATGTCGCAAGACGCCTCCAGCTTCTCGACCAGACCAG	è	1027 AlaproProAlaAspLysGluAlaPheA

laAlaGluAlaGlnLysLeuProGlyAsp 1045 ------AsnLeuGlnpro 1007 ::: CTGCGCATGTTCGACTTCACCG 5792 erProArgGlyLysSerArg---SerPro 1026 ||::: |cccgacgccrccrcaaarcacargcg 6032 CCCCGCACCGCACCCTCCACCC 5972 AACCGACCCACTCGACCAATCCACCTGCG 5912 | :::||| |---AGGAGTTCCTGCGCCATGCCTTCCC 6440 |||||||| SATCTTICCGTCGGCCGGTCCGGCCA 6380 ::: CTCGGCCAGTGCGTCGAGTGCGTGCAGGA 6320 CACCCCCACACCATCCGCACCCCCACCAA 6203 ----AlaProProAla----- 998 GCAGGGCGGTGACCCGGGCGGCGTCGTC 6730 CGATCTCGCCCTGCGAGTGCCCGGCCAC 6670 |||| |AAGGG-GGTGCGTCCGTGCCCCCCA 6551 ::: rrcgaggcrrggccagrcgacarcr 6491 AlaGluGlyAlaLeuLysAlaGluLysL 896 aLysserserGlyAlaProGlnAspSerA 916 euLeuSerProArgProSerLeuLeuThr 946 roglnLysPro----- 960 ysGlnArgAlaAlaAlaIleProProlle 976 luAspAlaAlaProThrLysPro---- 994 -----GlugiyProAlaL 876 ProProProAlaProProSerP 817 valLysserGluCysThrGluGluAlaG 871 GluProThrProAlaSerGluAlaThrG 804 GluGluGluThrAlaAlaAlaProProV 837 AlaGluGluLeuAla------

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3524 CCGGCGTCG---ACATCACCGTCACACCCCCGCCAACGCCAGCGAACACTCCCCCG---
                                     LeuSerLys-GluGluLeuIleGlnAsnMetAspArgValAspArgGluIleThrMetVa
                                                                       ----accrcaacectreacaceccagarecaececcaaceaceaceacea
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                                                                                                                                                                                                                                                                                                                                                           The present sequence represents DNA which encodes avermectin aglycon synthase proteins. Also described are: (1) polypeptides encoded by all or part of the DNA; (2) expression vectors containing the DNA; (3) host cells transformed by the vectors; (4) preparation of the polypeptides or culture of the transformatis; (5) preparation of avermectin aglycon or its derivatives by culture of transformed avermectin producing microorganisms; and (6) oligonuclectides of 5-60 bases in length containing sense or antisense sequences from the avermectin aglycon synthase DNA. The enzymes are useful for the production of modified forms of avermectin and of the intermediates in its biosynthesis, for use as drugs, veterinary drugs and agrochemicals
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methyltransferase from Streptomyces hygroscopicus that is
involved in the synthesis of the polyketide rappamicin"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      The present sequence contains a cluster of genes that encode polyketide synthases (PKSs) that are involved in the synthesis of soraphens in Sorangium cellulosum. The proteins encoded by the present sequence are SorR, SorA, SorB and SorB contain biosynthetic modules which contain a beta-ketoacylsynthase, an acyltransferase, a ketoreductase and an acyl carrier protein domain, as well as beta-ketone processing domains. S. cellulosum soraphens are useful as a cytostatic and antimicrobial agent active against phytopathogenic fungi. Soraphen-producing transgenic plants or biological control agents can also be produced, which may reduce crop losses and nutritional deprivation for
/note= "gene product is highly homologous to type I PKSs that are known to be involved in the synthesis of polyketide compounds" 942. .7115
                                                                                                                                                                                                                                                                                                                                                                                                            "gene product is highly homologous to type I PKS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   DNA encoding Sorangium cellulosum polypeptide(s) - used for, e.g.
biosyntheeis of soraphen useful as antimicrobial agent against
phytopathogenic fungi.
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                                                                                                                                                                                                                                                                                            "module 1 of SorB"
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/note= "module 2 of S
13455. .19616
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'note= "module 1 of
7203. .12884
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9871. .46318
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19871. .46318
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                                                                           misc_feature
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135

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22964 TCGACGCCGAAGGGTCGCA 1776Prol 23024 CTCCGCTCGCCGCCACCTC 1784		23384 AGCACGACGACCACCG 1861SerProlleSes ::: 23444 TCAGCGTCGCTCGCTCCCC 1878 uHisAsnThrGlyMetLys 23493	23564 GGAGCGCTCGGAGGCG 1927 YValTyrProThrLeumel 23624 CCCTCGTCCTCGGCCACG 1946 AArgProGluArgProArx 1962 SProProAraArgSerGl) 1962 SProProAlaArgSerGl) 23734 GCCACCGGCATCAAGCTC	23794 GCGCTCTTCATGCGCGAC 1983
8 8 8 8 8	6 8 6 8 6 8 6	8 8 8 8 8 8 8	6 6 6 6 6 6 6	4 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6 6
	1511ThralaSerSerSerGly 22101 TCTGACCACCGCCACCACGCCGACCTGGCCGGCACCTGGCTCGTCGTGGTGCGGGC 1520 GlySerIleAlaArgGlyAlaProValIleValProGluLeuGlyLySProArg	1578 LyaAlaSerGlnAspArgLysLeuThrSerThrProArgGluIleAlaLysSerProH18 15.320 GGGCTCCTCGCCCTCGACGAAGTCCCCTCGCCACCATGCCGCCGTGCCGC 15.98 SerThrValProGluHisHisProHisProIleSerProTyrGluHisLeuLeuArgGly	1643 LeuAspAlaAlaAlaAlaTyrTyrLeuProArgHis	1707 AlaAspMetLeuArgGlyLeuSerProArgGluSerSerLeuAlaLeuAsnTyrAlaAla 1726 1107 AlaAspMetLeuArgGlyLeuSerProArgGluSerSerLeuAlaLeuAsnTyrAlaAla 1726 1127 GlyProArgGrCGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGGG
8 8 8 8	8 4 8 4 8 4 8 6	86868686	8 8 8 8 8 8 8	888888

aaccagcgdcdcagrcrrddacagccggacacgrcgrggrgr 23263 erdly------GlydlyGlyGlyGerSerSerArgProAl 1852 GAAGGCCG-----CTTCCCAGCTCGACCCCGACAAGGCTTCT 23677 GATTCGATCATGACCGTCGAGCTTCGTCGGCGCTTGCAAC-AG 23733 CGTCGGCGATGTCGACGCTCTTTGGGAGTTCCTCCACCAAGGG 23973 31443 ||||| ccrccgagcacggcccggccccgaccrcgacaagcrcc 23563 GAGCAGCICCGTCTGCTCGCCGCTGGTGTGCGACGAGACGG 23623 AGGTCCGCGCCGTGTTCCACGCGGCGCCATCGGGCGCCACG 23023 ||||||| |CTCTCATGGAGGTCGCCGACGTTGTCTCTGCCAAGGTCCTAG 23083 SACCTGCTCGGTCCTCGACCCCTCGTCCTTTTCT 23143 lyLeuGluProAlaSerSer.---1974 euValProProValSerGlyHisAlaThrIleAlaArgThrPr 1998 roHisHisAlaSerProAspProProAlaPro----- 2014 ysGlyIleIleThrAlaValGluProSerLysProThrValLe 1898 erserProValArgProAlaAlaThrPheProProAlaThrHi 1918 etGluprovalLeuleuproLysGluAlapro---ArgvalAl 1946 rgAlaAsp-----ThrGlyHisAlaPheLeuAlaLy 1962 -----ArgProGlyThrGluGlnSerSe 1835 erHisAlaHisGlnHis----- 1860 erProArgThrGlnAspAlaLeuGlnGlnArgProSerValLe 1878 ------AspGl 1927 spargGluarqaspargAsparqGluarqGlubysSerIleie 1815 llGluHisAlaProlleTrp----- 1827 -----HisLeuThrLysProThrThrThrSerSerSerGlu 1795 oLeuSerProGlyGlyProThr hrLeu-----

1475 erLeuIleGlySerProGlyArg	δ	במסגס משפרפרנרנורפרפרנרנדשרפופשרופפונפרופנונפרפפופרונרפרפפיפר	
21870 CGCCTCCACGGCACCCGCCGCCGGCGTCAACCTTC	; A	Qy 1171 InLeuSerProArgGlyGlnAlaGlyProProGluSerLeuGlyValProThrAlaGlnG 1191	
1455 hrProLeulysTyrAspThrGlyAlaSerThrThrGl	8 8	Db 20866 GGGATGGCCCTCTCCCTGCTCGATACCTCGGCGCTCTTCCGGGCACAGCTCGAAGGGTGC 20925	
1443 roargProleuly: 21811 cccscsasaccscracccarccracccrrrc.	රු සි		
	g q	Qy 1139 uHisAlaLysAlaPro	
21703 GTGGTCGGCTCTCTGTGGAGCGACGAAGGGGATCTACC	g 2		
1406LysLeuLysProAlaHisGluGlyLeuVal	ò	1119 UArdCln1leGlyAla1leSerGlnGlyMetSerValGlnLeuHisValProTyrSerGl	_
21643 AGTCCTGTGCTGACCTTGCCCTTGCACGAGCTCCTCG	. q	Qy 1106 o	
	3 8		
1388	දු ද	Oy 1090 pThralaArgProValLeuProArgPro	-
	අ (Db 20575 GTGCTCGGCCAGGAGCGAGCCGGCCGTGCGCGCCCAGGCCCAGGCTCCGCGAC 20634	
1373 rgGluAlaLysLeuLeuLysArgGluGlyThrProPro	à i	1074 aPheSerTyrAlaproProGlyHisproLeuProLeuGlyLeuHisAs	_
21463 TGGCTCGGCGCTCCGACGGCGGTCCCGTTCTACT	QQ	Oy 1056ProvalProProArgGluvalIleLysAlaSerProHisAlaProAspProSerAl 1074	
1365 luAlaGln	ò	20458	
1353ThrGlr 1363GlySerileThrGlr 21403 TTGGCTGCAGGTGGAGTGGATTGGCG	දි සි	Qy 1042 uProGlyAapProProCysTrpThrSerGlyLeuProPhe 1055	•
21343 GCGGCCTGCTCCGCGATCTGGAGTCCGAGGGGGTCT	qq	Db 20413 TGGGCCGCCACGAAGATGGCAGTCCCGCGCGCGCGCGCGCGCGC	
1338 luArgHisSerProHisHisLe	ò	20356 AATCCCCTGATCGAGGATGCGCTCGCCATCGACGTCGATGGCACGAGGCG	
1318 aiglyArgAlaileSerSerAlaSerIleGluGlyLe 	දු පු	Qy 1004 nLeuGlnProGluSerAspAlaProGlnGlnProGlySerSerProArgGlyLy 1022	
21253 CGGCGCCTCCA	ପ୍ର	Qy 991 oThrLygProAlaProProAlaProProProProGlnAs 1004 Db 20296 AAGATCGTCGCCCTCCGCCATGACGCCTCCCCACCCTACACGCCCCGGCC 20355	
21193 GTGGAGCTCGCCGGCCAGGGGGCCATGGCCGCGGTGGT	දු ්	Db 20242 CTCAAGACCAACATTGGCCATCTGGGGGCCGCCTCGGGCCTCGCGGGCGTCGCC 20295	
1291 ys	λō	971 aAlaIleProProIleGlnValThrLysValHisGluProProArgGluAspAlaAlaPr	
21151 GAGGACGCTGCCAAGCTGGT	අ	Qy 954 nAlaSerProGlnLysProLeuAspLeuLysGlnLeuLysGlnArgAlaAl 971	
21103 AGCCAGGGCGAGATCGCCGCGGCCTCTCTCTT 11031 Indlv[vslvsGlvs]v	a &	Db 20131 GTCGAATGCCATGGCACCGGCACCTCCTTGGGAGACCCCATCGAGGTGCAA 20181	
erProSerArgLeuAspArgGlyArgGluAs	ò	934 sAsnArgLeuLeuSerProArgProSerLeuLeuThrProThrGlyAspProArgAlaAs	
21070 ATGGCGTCGAGCC	qq	20071 CAGAAAGTUCTCGGGGGGGCTCCAGAAAGGCCCTGCGAGAGGTC	
1231 isGlyThrProAlaAspValLeuTyrLysGlyThrIle	δ	DD ZOUII ACCGCCAIGAACCAIGACGGCGCGICGAGCGGCAICACCCCCCCAAAGGGCCACCCCCAC 20070 914 nSerbanSerSerblaThrCvqSerblabanGluValbanGlublaGluGlvAnGlvBanIv 934	
	අු	894 uLysGluGlyGlySerGlyArgAlaThrThrAlaLysSerSerGlyAlaProGlnAs	
lvileProSerThrArgValProS	3 8	Db 19951 CTCGAGCGCTCGGCGAGCCCCTCGCCCGAGGACACCGCGTCCTCGCCCTCGCGCGCC 20010	•
1191 luAlaSerValLeuArgGlyThrAlaLeuGlySerVal	8 8	Oy 874 oAlaLysGlyLysAspalaGluAlaAlaGluAlaThrAlaGluGlyAlaLeuLysAlaGl 894	

TGCTCCGCTCG-----AGGG 21920 cccccccccccccccccrc 21015 :: rcgcraccccraraccccrc' 21069 TGTGTGGCGCGCGCTGTCGCTC 21150 ::: ----GCGCTATGGCGATCGGCTC 21282 ATCTCCGGCGAGCCCCTGCCGTC 21342 CGCTTCTTCGTGGAGGTGAGCCCC 21642 CGGCGCTTCCTCGTCGCTCC 21762 ACGATCCT------GCCC 21810 C-AGCGCGAGCGCTTCTGGCTCGA 21869 -----cdAcccGGTGGTCGCCAT 21102 1.yMetSerValThrGlnCysSerL 1291 || ::: || ::: GTGGCGCGTGCGGCTC 21192 GAGCTGCCGGAGGCCGAGGTCGCA 21252 ||::: ||| TTCGCCCTCAAGCTGAGTTACGAC 21402 rccacegréadcescecesearé 21522 ||||||| | AACCTCCGCCAGCCGGTCCGCTTC 21582 hrTyrAspMetMetGluGlyArgV 1318 euMetGlyArgAlalleProProG 1338 InAlaLeuGlyProLeu----- 1405 isThrProGluLeuProLeuAlaP 1443 ysGluGlySerileThrGlnGlyT 1455 lySerLysLysHisAspValArgS 1475 ----GluAspGlyArgSerSers 1298 eutysGluGlnHisHisIleArg- 1352 -----GluAspTyrLeuArgA 1373 roProPro----- 1387 alalaThrValLysGlualaGlya 1423 rValProGlyGlySerIleThrLysG 1211 leThrTyrArgGlySerIleThrH 1231 leThrArgileileGlyGluAspS 1251 euProLysGlyHisVallleTyrG 1271 1391

18887			600 nserAlaGic 	620 tGluThrAla		640 tValGlyse	19069 ACCGGCATC	660 lnAsnLeuAs	19129 CTCTTCTTG	680 rgArgLysLy	19186 GACGCCGCC	700 ValGluAsp	19237 ATCGTCGGC	720	19297 TTCCTCGCC	731 lProArg		742 lAsnAsnSe	::: 19417 CTCGACCAG	755 sThrGluAl		774 yAlaAspGl	 19537 GGCATCGTC	785	19597 AGCGAATAC	796 oThrproAl	19657 ACCGCCGGG	808	19717 CTCTCGGTC	824 sGluGluLy	::: 19771 CAAGCCCTC	844 sProProAl	19831 TCCCCCGG	854 pThrGlyLy	19891 AAGACCTTC
qq	δ	අ ධ	Š	3 8	7 A	λõ	q	λō	q	ò	qa	ò	qa	à	ф	ò	: A	ò	: 음	ò	a	ð	අධ	ò	q	à	qa	ò	q _Q	ò	qq	ζ	셤	ò	Q
	275 aMethrativativationIlefienTvrPheLvsArdAshHisAlaArdLvsGlnTrpLy 295	croccecroscoscricargecccoaageaccarcercarcascaragra	295 sGlnLysPheCysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLy 312	18101 CCGGCGCCATTGGCGCTCACGTCGCCCGATGGCTCGCTCG			rTyrGluLysGlnPheProGluLleArgLysGlnArgGluLeuGlnGluArgMetGl	AGGAGCGCTGTCG	351 n SerArgValGlyGlnArg-GlySerGlyLeuSerMetSer 364 :::		365				395 leProProMetLeuTyrAspAlaAspGlnGlnArgIleLysPheIleAsnMetAsnGlyL 415 :::	18449 CTCGACCCTC 18459	415 euMetAlaAspProMetLysValTyrLysAspArgGlnValMetAsnMetTrpSerGluG 435	18460GACGCCTTTGTTCTCTTCTCGTCCGCGCGTCGTCTGGGGCGGCG 18505	435 InGluLysGluThrPheArgGluLysPheMetGlnHisProL 449	18506 GACAACAGGCGGCTATGCCGCTGCGAACGCCTTCCTCGATGCCCTGGCCGAGCAGCGGC 18565	449 ysAsnPheGlyLeulleAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuT 469	18566 GCAGCCTTGGGCTGACGGCGACATCGGTGGCCT 18598	469 yrTyrLyeuThrLysLysAsnGluAsnTyrLysSer	18599 GGGG-CGTGTGGGGGGGGGGGGGGTGGCTACGGGCTCCTGGCAGCCCAGCTAGAGCAA 18657	482	18658 cgcggrcrgrcgccgarggcccccrrcgcrggcgacgacgcrcgcr		18718 CACGACGACCACCCTCACCGTCGCCGACATCGACTGGGCGCGCTTTGCGCCTTCGTTC 18777		18778 AGCGCGCTCCCCGCCCCCCCCCCCCCGCGCATTTGCCCGAGGCGCAGCGCGCTCTCGAA 18837	525 LysGluLysGluAlaGluLysGluGluGlu-LysProGluValGluAsnAspLysGluAs 544		544 pLeuLeuLysGluLygThrAspAspThrSerGlyGluAspAsnAspGluLysGluAlaVa 564	18886	564 lAlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgArgLysGlyArglleThr 583
£	ì è	č, q	'n	qu	ò	qq	ò	qq	ò	셤	ò	අු	ò	qq	ò	qa	ò	đ	δλ	qa	δλ	qa	δ	QC	ò	đũ	ò	qg	ò	đ	ò	qa	ď	qa	ò

18887	ACAAGCTCCGAAACCGCTCGGAGAGCGAGAGAGAGAATCCACCTG 189:	1927
584	ArgSerMetalaAsnGluAlaAsnSerGluGluAlaIIeThrProGlnG1 600	10
600		20
62	tGluThrAlaLysLysGlyLeuLeuGluHis : : : TCGCTCATGACCGTCGAGCTTCGTC	9068
64	tValGlySerLysThrValSerGlnCysLysAsnPheTyr-PheAsnTyrLysLysArgG 6	60 9128
660	InAsnLeuAspGluIleLeuGlnGlnHisLysLeuLysMetGluLysGluArgAsnAlaA 68	30 9185
680	rgArgLysLysLys-LysAlaProAlaAlaAlaSerGluGluAlaAlaPheProProVal 69	99 9236
19237	ValGluAspGluGluMetGluAlaSerGlyValSerGlyAsnGluGluGluGluGluGluGlu 71	19 9296
720	Trectedectargeaceceargecearteceargeceratecers 13	31 9356
731	IProArg	42 9416
742	AshAshSerSerAspThrGluSerIleProSer	55 9476
755	### STHE GLUALAALALYSABPTHEGlyGlnAsnGlyProLysProProAlaThrLeuGl 77	74 9536
77	yalaAspGlyProProProGlyProProThr78	84 9596
78	AGCGAATACGCACTGCGAACACGAGGACGAAGGTCGAAGGCGTATGCCCTCCAAGGC 19	96 9656
796	oThrProAlaSerGluAlaThrGlyAlaProThr	
19657	Accecedercerreceseseseseseses 1	
808	ProProProAlaProProSerProSerAlaProProProValValProLy 82	9770
824	sGluGluLysGluGluGluThrAlaAlaAlaProProValGluGluGluGluGlnLy 84.	9830
844		854 19890
85	pThrGlyLysAlaGluGluProValLysSerGluCysThrGluGluAlaGluGluGluGlyPr 	874 19950

us-09-522-753-5.rng

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pyrrolnitrin. Pyrrolnitrin has antibiotic activity, and inhibits a broad range of fungi. It is a fungal respiratory electron transport inhibitor. The genes are useful for developing a host as a biocontrol agent, for
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   chlorotryptophan to monodechloroaminopyrrolnitrin. PrnC enzyme catalyses the conversion of monodechloroaminopyrrolnitrin to aminopyrrolnitrin. PrnD enzyme catalyses the conversion of aminopyrrolnitrin to
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         protecting plants against a pathogen and for producing large, uniform amounts of pyrrolnitrin. The genes are also useful for producing transgenic plants that express antipathogenic substance (APS) biosynthetic genes. The genes and enzymes are useful for increasing the resistance of host plants to disease caused by phytopathogens, e.g. fungl, bacteria or nematrodes. The microbial hosts are useful for controlling or inhibiting the growth of a phytopathogen. (Updated on 15-2003 to standardise OS field)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The present sequence represents a Sorangium cellulosum soraphen gene cluster. The specification describes pyzrolnitrin gene regions. This region encodes enzymes required in the biosynthetic pathway of pyzrolnitrin. The PrnA enzyme catalyses the conversion of tryptophan to chlorotryptophan. The PrnB enzyme catalyses the conversion of 7
                                                       2473 laGlyValMetAlaSerProProProProGlyLeuProAlaGlySerGlyProLeuAlaG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Pyrrolnitrin biosynthesis genes and enzymes, useful for producing pyrrolnitrin for increasing plant resistance to phytopathogens, e.g. fungi or nematodes, and for developing microbial hosts useful as
                                                                                                                                                                                                                                                                                                                                                                        Pyrrolnitrin, PrnA, PrnB, PrnC, PrnD; antibiotic, biocontrol, fungal respiratory electron transport inhibitor; transgenic plant; antipathogenic substance; biosynthetic gene; fungal resistance;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 TR;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Young
                                                                                                                                                                                                                                                                                                                                     sequence of Sorangium cellulosum soraphen gene region.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Kirner
                                                                                                                                        25389 AGCCCACAACCGCGACGAGCTCCTCCGCGC 25420
                                                                                                       2493 lyProHisHisAlaTrpAspGluGluProLys 2503
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Van Pee K,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Example 14; Col 121-146; 124pp; English
                                                                                                                                                                                                              AAA75299 standard; DNA; 28958 BP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        94US-00258261.
95WO-IB000414.
96US-00729214.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              (NOVS ) NOVARTIS FINANCE CORP
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        98US-00028934
                                                                                                                                                                                                                                                                                                                                                                                                                               phytopathogen; soraphen; ss.
                                                                                                                                                                                                                                                                                                  (first entry)
                                                                                                                                                                                                                                                                                   (revised)
                                                                                                                                                                                                                                                                                                                                                                                                                                                               Polyangium cellulosum
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      WPI; 2000-593649/56.
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505	InGinGinGinGinGinProMetPro-ArgSerSerGinGiuGiuLy8AspGiuLy8Giu 	524
525	LysGluLysGluAlaGluLysGluGluGlu-LysProGluValGluAsnAspLysGluAs	544
18838	GCCAGCGCCGATGCGTCCTCCGAGCAGAGGCCACAGGCCTCCTCG	18886
544	pleuleulysGlulysThrAspAspThrSerGlyGluAspAsnAspGluLysGluAlaVa 56	64
18886	18	18886
564	lAlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgArgLysGlyArglleThr 58	
18887	ACAAGCTCCGAAACGCTCGGAGAGCGAGAAGAGCAAGATCCACCTG	8927
584	ArgSerMetAlaAsnGluAlaAsnSerGluGluAlaIleThrProGlnGl 	600
600		20
18988	::	9026
19027	tGluThrAlaLysLysGlyLeuLeuGluHisGlyArgAsnTrpSerAlaIleAlaArgMe 6 : :::	640
64	tValGlySerLysThrValSerGlnCysLysAsnPheTyr-PheAsnTyrLysLysArgG 6	. 09
9	ACGGGATCAAGCTCCCGGCCACCCTCGACCATCCCTCTCCTCAACGGCGTCGCG	19128
099	InAsnLeuAspGluIleLeuGlnGlnHisLysLeuLysMetGluLysGluArgAsnAlaA	680
19129	crcrrcrrcccccacrcccrcccccccccccccccc	19185
680	rgArglysLysLys-lysAlaProAlaAlaAlaSerGluGluAlaAlaPheProProVal	699
700	ValGluAspGluGluMetGluAlaSerGlyValSerGlyAsnGluGluGluMetValGlu	719
19237		9536
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17348 TGCTCCCTCCCACCTCTCGAGCTTTCACCACGCGCCCAAGAGCAGGACCACGGTCGACA 17407
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                                                                                                                                                                                                                                                                  Protecting plants against pathogens with genetically transformed biological control agent - which expresses all polypeptide(s) involved in pyrrolnitrin biosynthetic pathway.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                SerThrGlnLeuValAlaGlnThrTrpArgAlaThrGluProArgTyrProProHisSer
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92US-00937648.
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Polyangium cellulosum
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Best Local Similarity:
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transgenic plant; phytopathogen; resistance; ss.
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                                                                                                                                                  rnis is the soraphen gene cluster from S. cellulosum. The gene cluster may be expressed recombinantly to produce soraphen, or expressed in a transgenic plant for disease-resistance. (Updated on 16-OCT-2003 to standardise OS field)
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                                                                   New genes for biosynthesis of anti-pathogenic substances - pref.
pyrrolnitrin and soraphen, useful for disease control in plants.
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අ <u>ග</u>	CGGCTTGCGCGCCGCCGCACTGTGTGTGCCGCCTCACGTTTCACGTGTGTCTCTATATTTCCCTG	ò	
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qq	9079 TCTATICTIGGCACCGCTCGGAGGGGACCGGTGCGCGCTGCCGTTTATCCCGGCGGACG 9138	ò	
ò	414 lyLeuMetAlaAspProMetLysValTyrLysAspArgGlnValMetAsnMetTrpSerG 434		2
qq	9139 TCATGATGTCC 9149	ò	
ò	434 luGlnGluLysGluThrPheArgGluLysPheMetGlnHisProLysAsnPheGlyLeuI 454		10
Q	9150ACCGGGAGGAAACGTGGAAATC-AAGTACATGGGTCAG 9187		
ò	454 leAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyrTyrTyrLeuThrL 474		10
QQ	9188TTGACCA 9194	ò	
à	474 ysLysAsnGluAsnTyr-LysSerLeuValArgArgSerTyrArg-ArgArgGlyLysSe 493		12
ф	9195 recedracadecececadadecrecerecencececedececadececececece 9254	ò	
'n	493 rGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnG		2
Db	9255 TTCTGCTCCTCAACGACAAGACACCCGTCACGACCACCCAC	ò	
à	513 oArgSerSerGlnGluGluLysAspGluLysGluLysGluLysGluLysGluLysGluLysGluGl 533		10
Db	9315 GCGAGARACCCGCCGCGCGCGCCCTGACCTCCAGACCTATATTCTCCCAGCTCCGGA 9374		
ò	533 uGluLysProGluValGluAsnAspLysGluAspLeu-LeuLysGluLysThr 550		-
Ωp	9375 AATGCCTCGCGGCGATGAGCGGGGGAAGCCTCGCCTGCTATCCGAGAAAACCCTGGTGA 9434		i
ò	551		,
යි	9435 CGTGGCCCTGCGGTTATCTGGCGCGCGCGCGCGCGCCCCTTCGGATGTGG 9488		í
ò	559 spGluLysGluAlaValAlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgArgL 579	÷ 6	,
qq	9489 CTGAATTCCGTCGGTTTGCCAGGGGGGGGGGAAGCGGAACGCA 9533		á
ò	579 ysGlyArglleThrArgSerMetAlaAsnGluAlaAsnSerGluGluAlaIleThrProG 599		
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Q C	GCCAGGGACCACTCCTCGCCGATATCGAGCACGGCTGCTGCGCGCCGCCGAGGCGTAC	ò	
ò		ਦ ਰੂ	H
qq	9645 GCATGGAGGAGTGCCGGCTTTCGCTCGTCGAACGCTCCATCGAGGGGGGGCGACCTGCTCG 9704	ò	
δ	631 lyArgAsnTrpSerAlalleAlaArgMetValGlySerLysThrValSerGlnCysLysA 651		ä
qq	9705 GCGGCACCGGGAGGTCGTCAGCGAACTGTCCGCCCTCGTCGCCCAG 9751	ò	
ò	651 snPheTyrPheAsnTyrLysLysArgGlnAsnLeuAspGluIleLeuGlnGlnHisLysL 671		Ä
qq	9752TACCCCTACCACGAACAGCTCACGGGCAGCTCATGGTCGCGCTCG 9797		
ò	671 euLysMetGluLysGluArgAsnAlaArgArgLysLysLysLysAlaProAlaAlaAlaS 691		_
qq	9798 TGCGCTGTGGCCGCCGCGGGAGGCGCTCGCGTGCACCAACGGCTCCGGGCCCGCATGG 9857)
ò	691 erGluGluAlaAlaPheProProValValGluAspGluGluMetGluAlaSerGlyValS 711		Ä
qa	9858 TGGAGGACCTCGGCCTCGAACCGAGCAGCCACCTGCGCGCCCTCCTGCCGCCGTCCTCA 9917		ř.

804 1yAlabroThrProProProAlaProProSerProSerAlaProProProValValProL 824 10903 ::::::|| || || || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 || 1000 | 619 CACGGGAACTCCACCTGGGCCGCGTAGTTGAGCCCCGCGCCGAACCCCCACGAGGAGC 10678 1499 GICCICCICGCGCGICGICACGAGGACCACCCCGCGGGGAAGCACGACATCGCGGCGCG 10558 020 CGGGGCCCGGTCCCCGTCGCCGGGTCGAAGGCCTGGACGGAAAAAGGCCCT 10079 080 CICCIICCCGCCGCCGGGGCCCGCGCTCAGCGCGCGCGAAACCCCCGCTCCCAGTGC 10139 140 ACGAGCGGTTCGCCCTCACCCTGCCCCACCC-CGCGCACCTCGCCGACGACGATCACGTG 10198 ||||||::||| 199 GTCCCGGCCAGGTGCCGGGCGTGTACCGCGCACTCCAGCGTGACGAGGCGCGTCCGCCGC 10258 918 GCGGCCCGGAACCGGGACCGCTGCCGGGACCGCCCGGAACCGGTGGCGAGA 9959 780 ----- 782 804 ----- 804 824 ysGluGluLysGluGlu---GluThrAlaAlaAlaProProValGluGluGlyGluGluG 843 lnLysProproAlaAlaGluGluLeuAlaValAspThrGlyLysAlaGluGluProValL 863 863 ysSerGlu-CysThrGluGluAlaGluGluGlyProAlaLys------ 876 884 -------GluAlaThrAlaGluGlyAlaLeuLysAlaGluLysLysGluGly 898 899 GlySerGlyArgAlaThrThrAla-LysSerSerGlyAlaProGlnAspSerAspSerSe 918 918 ralaThr----CysSerAlaAspGluValAspGluAlaGluGlyGlyAspLysAsnArgLe 937 753 er----- 753 766 lyprolyspro------proAlaT 772 711 erglyasnGluGluGluMetValGluGluAlaGluAlaLeuHisAlaSerGlyAsnGluV 731 731 alproArgGlyGluCysSerGlyProAlaThrValAsnAsnSerSerAsp----------ThrGluSerIleProS roAlaSerGlu---AlaThr------798

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                                         60 GlnProGlnArgArgArgProSerLeuLeuSerGluPheGlnProGlyAsnGluArgSer
                                                                                80 GlnGluLeuHisLeuArgProGluSerHisSerTyrLeuProGluLeuGlyLysSerGlu
                                                                                                                                                                                                       100 MetGluPhelleGluSerLysArgProArgLeuGlu-----LeuLeuProAspProLeu
                                                                                                                                                                                                                                                3201 GGTGGTGGCGATGACCAGCGGCGCGCTCTTCAGCATGTTCTACTTCCTCACCCTCTT
                                                                                                                                                                                                                                                                                                                                8261 CCTCAACCAGGTACGGGACTACAGCCCGCTGCGCACCGGCT-TCGCCTATCTGCCGCTCG
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8060 CTGGGGCTCCGCCGCCACCCTCGTCCC-
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            The present sequence represents a frenolicin gene cluster from Streptomyces roseofulvus. The present invention describes: (1) a vector containing the frenolicin gene cluster DNA sequence operably connected to an expression control sequence; (2) a host cell transformed by the above the preparation of frenolicin or a biosynthetic intermediate for it in which the above cell is cultured and frenolicin or its biosynthetic intermediate for it in which the above cell is cultured and frenolicin or its biosynthetic intermediate is isolated from the culture or the cell; (5) a method for intermediate is isolated from the culture or the cell; (6) a method for the preparation of frenolicin B by axidising frenolicin, and (6) a method for the preparation of a feed composition by mixing frenolicin with other components. Frenolicin B is useful as an antibiotic
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         14 AlaThrGluProArgTyrProProHisSerLeuSerTyrProValGlnIleAlaArgThr 33
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AAWS5806, AAWS5807, AAWS5808, AAWS5809, AAWS5810, AAWS5811, AAWS5812,
AAWS5813, AAWS5814, AAWS5815, AAWS5816, AAWS5817, AAWS5818, AAWS5819.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Sequence 24379 BP; 3077 A; 9792 C; 8499 G; 3011 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                               /transl except= (pos:22040. .22038,aa:Arg)
/transl except= (pos:22037. .22035,aa:Arg)
/transl except= (pos:22034. .22032,aa:Thr)
/transl except= (pos:22031. .22029,aa:Lys)
/note= "encodes protein given in AAW55819"
                                                                                                     /*tag= r
/note= "encodes protein given in AAW55817"
complement(19990. .20907)
                                                                                                                                                                                 /transl_except= (pos:20907. .20905,aa:Met)
/note= "encodes protein given in AAW55818"
"encodes protein given in AAW55815'
                                                            AAW55816
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688
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                                                            "encodes protein given in
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Matches:
Conservative:
Mismatches:
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/note= "encodes protein given in AAW55806"
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complement (10618. .11628)
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/transl_except= (pos:11628. .11626,aa:Met)
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complement(16453. .16935)
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complement(12154. .13209)
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                   2237 SerArgSerAlaValTyrProLeuLeuTyrArgAspGlyGluGlnThrGluProSerArg
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8 8 8	 13661 CTCGCCGACAGCGGAC	g & g	14471 GAGATCGCGGCCTGGCGCTCGCCATGGAGTACGGCGTGGTGCCGCCCACCGCCAACCTC 14530 1931 hrLeumetGluProValLeuLeuProLysGluAlaProArgValAlaArg 1947
ර් සි	aTyrTyrLeuProArgHisLeuAlaProAsnProThrTyrProHisLeuTyrProProTy	s d	1948ProA 1952 14591 GTGGACAGGGTCCTGAGCGTCGGCGGCTTCGGCGGTTTCCAGAGCGCCTTGGTCCTC 14650
8 & B	ProAspThralaAlaLeuGluAsnArgGlnThrileIleAs	yo da	
상 원		oy O	ProAlaArgSerGlyLeuGluProAlaSerSerProSerLysGlySerGluProArg
දි සි	1707 laaspMetLeuArgGlyLeuSerProArgGluSerSerLeuAlaLeuAsnTyrAlaAlaG 1727 	Qy Qp	
& 8	1727 lyProArgGlyIleIleAspLeuSerGlnValProHisLeuProValLeuValProProT 1747 13905	Sp GS	2003 AlaProHisHisAlaSerProAspProProAlaProProAlaSerAlaSerAspProHis 2022
8 8	hrProGlyThrProAlaThrAlaMetAspArgLeuAlaTyrLeuProThr	පු ද	2023 ArgGluLysThrGlnSerLysProPheSerIleGlnGluLeuGluLeuArgSerLeuGly 2042
ે ઠે ક	SerP	oy Ob	2043 TyrHisGlySerSerTyrSerProGluGlyValGluProValSerProValSerPro 2062
3 6 8		රු අු	2063 SerLeuThrHisAspLysGlyLeuProLysHisLeuGluGluLeuAspLysSerHis 2081
8 8 8		රු අ	2082 LeudludlyGluLeuargProLy8
3 &	19088 CCITCGACAGGGGAACGGGGTTCGTCCTCCTCCTCTCTCTCTC	& 6	2090
da V	14122GGAĠĠĠCGCCGCC 14134 1839 lySerSerGlyGlyGlyGlySerSerArgPro	B & 1	AlaalatisLeuProttisLeukrgProLeuProGluSerGlnProSerSerProLeu
දු දු	GTCCTCGTGCTGGAGGAGCTGGGGCACGCGGGGCCCGCGACGTCTACGCTCTCAlaSerHisSerHisAlaHis	<u>ک</u> ۾	2120 LeuGlnThralaProGlyValLysGlyHisGlnArgValValThrLeuAla 2136
8 8 8	14195 GIGTCCGGCTACGCGTGCAACGCGTACCATGACCGGGCTCACCGCGCGCG	oy . Db	2137 GlnHisIleSerGluValIleThrGlnAspTyrThrArgHisHisProGlnGlnLeuSer 2156
8 8		oy Pp Q	2157 AlaProLeuProAlaProLeuTyrSerPheProGlyAlaSerCysProValLeuAspLeu 2176
8 6	avalgluProSerLysProThrValLeuArgSerThrSerThrSerSerProValArgP	cy B	ArgArgProProSerAspLeuTyrLeuProProProAspHisGlyAlaProAlaArgGly
ර ය	1909 roAlaAlaThrPheProProAlaThrHisCysProLeuGly 1922 	6 G	SerProHisSerGludlyGlyLygskrgSerProGluProAsnLygfhrSerValLeuGly 221 ::: :: GCTGGCGGTGAGCGGCGGCGTGGAGTCCGCGCTCGACCCCTGGG 154
خ	1923GlyThrLeuAspGlyValTyrProf 1931	S Q	2217 GlyGlyGluAspGlyIleGluProValSerProProFroGluGlyMetThrGluBroGlyHis 2236 2217 GlyGlyGluAspGlyIleGluProValSerProProFroGluGlyMetThrGluBroGlyHis 2236 35402 CCTGGCCGCCCACGCCTCGTCGGGCACCCTCAGCCGCTCCGGCGACCCGGCCAC 15455

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GOGGGGCCCCGGGCCCGGCCCGGCCGGCGCGCGCCGCCCCCGCG	11417 ĠĊĠCCGGCCCCATGAACCGCAAGGTCTCGTCCGGCGGCGAAGCCGCGCGCG
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us-09-522-753-5.rng

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This DNA sequence comprises the Streptomyces frenolicin gene cluster containing specifically claimed coding sequences (genes A-U) that respectively encode 21 proteins (see AAM14199-219) involved in frenolicin synthaesis. The genes can be divided into 5 subclusters: (1) genes A, B, C, D and F encode an efflux pump; (2) genes H, I, J and K encode butyrate starter synthaese; (3) genes L, M and N encode polyketide synthaese (PKS); (4) genes O, P, Q and R encode a hemiketalase, a ketoreductase and cyclases/dehydrases; and (5) genes S and T encode a keto/enoyl reductase and a hydrolase. Also claimed are vectors, host cells (especially a Streptomyces pp., particularly Streptomyces roseofulvus), and the encoded proteins. Cells transformed using the above sequence can be cultured to
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Streptomyces frenolicin gene cluster - useful for producing recombinant frenolicin antibiotics.
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AAW34205, AAW34206, AAW34207, AAW34208, AAW34209, AAW34210, AAW34211,
AAW34212, AAW34213, AAW34214, AAW34215, AAW34216, AAW34217, AAW34218,
                                                                                                                                                                                                                                                                                                                                                      'product= "cyclase/dehydrase related to act VII"
'note= "gene Q (specifically claimed)"
18895. .19839
                                                                                                                                       'product= "acyl carrier protein used by the PKS"
'note= "gene N (specifically claimed)"
                                                                                                                                                                                                                                                                                                                                                                                                                         related to actIV" claimed)"
                                                                                                                                                                                                                                                               *tag= p productase related to actIII"
'product= "ketoreductase related)"
'note= "gene" (specifically claimed)"
                                                                                                                                                                                              *tag= o
product= "putative hemiketal dehydrase"
product= "PKS ketoacylsynthase subunit"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      /product= "quinone-forming hydroxylase"
/note= "gene T (specifically claimed)"
complement(22505. .22179)
                                                   ^*tag= m
/product= "PKS chain length factor"
note= "gene M (specifically claimed)"
16120. .16371
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    /note= "gene S (specifically claimed)"
complement(20904. .22094)
                L (specifically claimed)"
                                                                                                                                                                                                                              note= "gene O (specifically claimed)".7088. .17903
                                                                                                                                                                                                                                                                                                                                                                                                                           /product= "cyclase/dehydrase
/note= "gene R (specifically
complement(19990. .20907)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                'product= "oxidoreductase"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           /product= "not specified"
/note= "gene U"
                                                                                                                                                                              complement (16453. .16935)
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                "gene L
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                                                                                                                          *tag=
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ThriyaAspArgSerLeuThrGlyLysLeuGluProValSerProProSerProProHis 155
                                                  305 ACCAAGGACCGTAGCCTGACGGCAAGCTGGAACCGGTGTCTCCCCCCAGCCCCCGCAC 364
                                                                                                                                                                                                                                                                                                                                    Frenolicin; antibiotic; feed additive; anticoccidial; coccidiostatic; efflux pump; butyrate starter synthase; polyketide synthase; PKS; hemiketalase; ketoreductase; cyclase; dehydrase; ketoreductase; hydroxylase; Streptomyces roseofulvus; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               /product= "ATP-binding component of ABC transporter"
/note= "gene D (specifically claimed)"
5533. .7183
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           'product= "protein with 6 membrane-spanning domains"
note= "gene C (specifically claimed)"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        /product= "translationally coupled to gene I"
/note= "gene H (specifically claimed)"
complement(10618. .11628)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        'product= "unknown non-membrane protein"
note= "gene E (specifically claimed)"
344. .8897
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       /product= "80 kDa non-membrane protein"
/note= "gene A (specifically claimed)"
2945. .3916
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             /product= "putative efflux pump"
/note= "gene F (specifically claimed)"
9164. :10012
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 /product= "membrane protein"
/note= "gene B (specifically claimed)"
4020. .4844
/*tag= c
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/product= "transcription activator"
/note= "gene G (specifically claimed)"
complement(10105. .10621)
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/product= "acyltransferase"
/note= "gene K (specifically claimed)"
13409. .14686
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/note= "gene I (specifically claimed)
11809. ,12066
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/note= "gene J (specifically claimed)
complement(12154. .13209)
                                                                                                                                                                                                                                                                                                    Streptomyces frenolicin gene cluster.
                                                                                                                                                                                                                                                                                                                                                                                                                                                              Location/Qualifiers
                                                                                                                                                                                              AAT93095 standard; cDNA; 24379 BP
                                                                                       ThraspProGluLeu 160
                                                                                                                      365 Acreacerreagerie 379
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Jones LW;

Dickson MC,

Stache-Crain B,

Drmanac RT, Labat I,

(JONE/)

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STACHE-CRAIN DICKSON M C. JONES L W.

30-JUL-2001; 2001US-00918995 30-JUL-2001; 2001US-00918995

17-APR-2003

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CCGGTGCCGCCGCCGCCGCTGCCCGTTGGTAGCTGAGGAGGCTGCGC-CGGCGCC 43076
                         -----TGTCCACCGGAGCCGGCGGGA 43473
                                                                                                                                                                                                                                                                                               ------CCGGTGCCGCCTTGGCCGC 43368
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                                                                                                        snSerAlaMetValLysSerLysLysGlnGluIleAsnLysLysLeuAsnThrHisAsnA 2299
erAlaValTyrProLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlyS 2259
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mapping; biodiversity; genetic disorder.
                                                                                                                                                            2299 rgAsnGluProGluTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetProAlaI
                                                                                                                                                                                                                 2319 leThrGlyThrGlyLeuMetThrTyrArgSerGlnAlaValGlnGluHisAlaSerThrA
                                                                                                                                                                                                                                                                    2339 snMetGlyLeuGluAlaIleIleArgLysAlaLeuMetGlyLysTyrAspGlnTrpGluG
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                         CAGCGTTACCGCCT---
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The invention relates to an isolated polymuclectide comprising any one of 38043 cDNA sequences, appearing as ACH12789-ACH50831, whose sequence was determined by the technique of SBH (sequencing by hybridisation). Also included is a purified polypeptide comprising a sequence corresponding to a reading frame of the novel polymoleotide. The nucleic acid sequences are useful in diagnostics as expressed sequence tags (EST) for informatics, in assessing biodiversities, or in identifying mutations in forensics, in assessing biodiversities, or in identifying mutations responsible for genetic disorders and other traits. The nucleotide for corresponding in the recombinant production of protein, or in generating antibodies specific for it. The present sequence is useful for generating antibodies specific for it. The present sequence is one of the 38043 isolated cDNA/EST sequences. Note: The sequence data for this patent did not form part of the printed specification, but was obtained in electronic format directly from 18770 at
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 124
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                                                                                                                                                                                                                                                                                      New polynucleotide sequences obtained from various cDNA libraries, used as hybridization probes, as oligomers for PCR, for chromosome and gene mapping, in the recombinant production of protein, or in generating
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CCCCTGCTGCGACCGTCACCCCTGCTGGACACGGGCCAACCTGCGGGATCTGAAGACCTC

GlyLysSerGluMetGluPheIleGluSerLysArgProArgLeuGluLeuLeuProAsp 115

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	257TyrAsnGlnProSerAspTh 48629 CCGGTGCCCGGTGCCGCGTTGAAGTTGGCCCCGCCGCCGCCGCCGCCGCCGCCGCCGCCGCCG	282 uTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPheCysGlnArgTy 282 uTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPheCysGlnArgTy 28509 GCGTTGCCGCCCCCGGTGCCGCCCCCCGGTGCCGCCGGTGCCGCC	48449 CIGCCGTIGCTGAAGCCGACCCCCGGCGACGACGACCACCCGCGCGCG	362 rMetSerAlaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSerGluGl 	aAspGl	442 uLysPheMetGlnHisProLysAsnPheGlyLeuIleAlaSerPheLeuGluArgLysTh ::: ::: ::: ::: 48176 CCGGCCGCTCCGGCCGCTCCGGCCGCTCCGCCGGTGCCGCC	48116 GTGCCGCCGCATGCCGGTGGGGTTATCGGCGCCCGGCCGG	47996 CCGGCCCGCGACACCGGCGTTG-CCTGTGGTGCCGATCATGCCGCCGCGCTCCGGT 517 nGluGluLysAspGluLysGluLysGluLysGluAlaGluLysGluGluGluGluLysProGl 47937 GCCGGCCGCCGGCCGGCCGGCCGCTCGGCCGCTCGGTGCCGCTCGCCGC		

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CGGTGGGTTATCGGCGCCGCCC------ 4748 coArgGlyGluCysSerGlyProAlaThrValAsn 743 SluGluThrAlaAlaAlaProProValGluGluGl 840 sluGluLeuAlayajAspThrGjyLysAjaGluGl 860 ijyAlaLeuLysAlaGluLysLysGluGlyGlySe 900 laGluGlyGlyAspLysAsnArgLeuLeuSer-P 940 ThralaLysLysGlyLeuLeuGluHisGlyArgA 633 GlyserLysThrVal------646 MetGluLysGluArgAsnAlaArgArgLysLysL 684 uGluAlaAlaPheProProValValGluAspGlu 703 yAsnGluGluGluMetValGluGluAlaGluAla 723 roSerProHisThrGluAlaAlaLy8AspThrGl 763 ThrLeuGlyAlaAspGlyPro----ProProGl 781 serArgAlaProlleGluProThrProAlaSerGl 801 AlaGluLeuAlaSerMetGluLeuAsnGluSerS 613

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Qy 2317Thr 232 Db 85205 ACCACCACGCTTCGCCTACGCCCCGCTTCCAGGCCTGCGGGCCTGGCGCCGCGCGCG	Oy 2360SerProProLeuSerAlaAsnPheAsnProLeuAsnAlaSerLeu 2377 Db 85445 GCGCTACGCGTCGGCTCGCCCGAACGGCCTGTCCGTCACCGCCGAC 85504 Oy 2378 ProAlaAlaMetProIleThrAlaAlaAspGlyArgSerAspHisThrLeuThrSerPro 2397	85612 GGACTGGA 2436 GluGlyA8 8572 CCCGGACA 2447 ABRARGVA 85732 CCTCGA 2467 Leu	2482	RESULT 49 AA199682_39/C Continuation (40 of 45) of AA199682 from base 3900001 (Mycobacterium tuberculosis strain WP Sequence split into 45 fragments LOCUS AA199682 Accession Aai99682 WP Fragment Name Begin End WP AA199682_01 100001 210000 WP AA199682_03 300001 310000 WP AA199682_04 400001 310000 WP AA199682_05 500001 310000 WP AA199682_06 600001 710000 WP AA199682_07 700001 810000 WP AA199682_09 900001 1010000 WP AA199682_10 1100001 1110000 WP AA199682_11 1100001 1310000 WP AA199682_12 1200001 1310000 WP AA199682_13 1300001 1510000 WP AA199682_13 1300001 1510000 WP AA199682_13 1300001 1510000 WP AA199682_13 1300001 1610000 WP AA199682_14 1400001 1510000

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ራ -	1354 rIleThrGlnGlyIleProArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgG1 1374	Db 82156 CAGCCAGGGCG
ò	uAlaLysLeuLeuLysArgGluGlyThrProPr	82203
QQ		Qy 1744 alProProThr
රි ස්	1394 rGlualaTyrLysThrGlnalaLeuGlyProLeuLysEcuLysProhlaHisGluGlyLe 1414 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	82246
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ò	1433beuArgHisThrProGluLeuProLeuAlaProArgProLeuLysGluGl 1449	DD 82366 ACTCTCTGGG
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۵	82426	CACCGGCGTCTTCGCCGGCGTCATGTACAACGACTACGGCACCCCTCTCTCGCCGGCGACGA 82485
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۵	82486	GTACGAGGCGTTCCGCGGCAACGGCAGCGCCCCGAGCGTCGCCTCCGGCCGCGTCTCCTA
>-	1823	HisAlaProlleTrpArgProGlyThr
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1941 GGGCGGGCGCGCCCCCTACCCCTTCCAGCACCGGGGGTTCTGGCCGCACCGGGGGC 119 - GluGluAlaGluAlaCutianianias and processor 119 - GluGluAlaGluAlaCutianianias 119 - GluGluAlaGluAlaCutianianias 110 - GluGluAlaGluAlaCutianianias 110 - GluGluAlaGluAlaCutianianias 110 - GluGluAlaGluAlaCutianianias 110 - GluGluAlaGluAlaCutianianias 110 - GluGluAlaGluAlaCutianianias 110 - GluGluAlaCutianianias 110 - GluGluAlaCutianianias 110 - GluGluAlaCutianiania 110 - GluGluAlaCutianiania 110 - GluGluAlaCutianianiania 110 - GluGluAlaCutianianianianianianianianianianianianiani	80218 915 935 80338 80389 975 80421
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1278	B lyArgGluAspSerLeuProLysGlyHisVallleTyrGluGlyLysLysGlyHisValL	1258 81173
1258	<pre>B euTyrLysGlyThrIleThrArgIleIleGlyGluAspSerProSerArgLeuAspArgG</pre>	1238
1238		1223
1222	5 lyGlySerIleThrLysGlyIleProSerThrArgValProSerAspSerAla	1205
1205	7ProThrAlaGlnGluAlaSerValLeuArgGlyThrAlaLeuGlySerValProG	1187
1186 80961	5ArgGlyGlnAlaGlyProProGluSerLeuGlyVal	1175
1174	6 etApproLysLysLeuAlaProPheSerGlyValLysGlnGluGlnLeuSerPro	1156
1156	6 rolyrSerGluHisAlaLysAlaProValGlyProValThrMetGlyLeuProLeuProm	1136 80782
1136 8078]	6 erVälLeuGluArgGlnIleGlyAlaIleSerGlnGlyMetSerValGlnLeuHisValP 	1116 80773
1116	6 euProArgProProThrIleSerAanProProProLeuIleSerSerAlaLysHisProS	1096
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694	78234 CGTGGGGGCTCTCCGCCGCCGCCGCCGCCCTCGACGCCCGCC	q
682 LYSLYSLYSLYSALA-FIGHTAN 		ò
79172 GICCGCTICG	33	<u></u>
662		යු ,
79112 GTCGCCGTCGCCGACCTGTGCTCCGCC	356 lnArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluHisGluValSerGluIlel 376	ò
643SerLysThrValSerGlnCysLysAsr		ద
79052 GCCGCGGCCTGACCTACCACGAGCCGCGC	LysGlnArgGluLeuGlnGluArgMetGln-SerArgValGlyG 356	ò
, 624 LysLysglyLeuLeuGluHisGlyArgAsn	78006 TGCCGCGCAGCCTGCTACCCTACCCACCACCACGACCTCGACTGGACCGCCGGCTCCG 78065 Qy	g
79031	337	à
604	77970 TGATCAAAATGATCATGGCGTTGCAGCGCGG	d d
78992	302 rAspGlnLeuMetGluAlaLeuGluLysLysValGluArglleGluAsnAsnProArgAr 322 Db	ò
	77959Tgccccddcg 77969	qq
78935	302	ò
564	77946 AGGCCGC	g
10000	282	ò
544 Asn[.e.]	77945	q
78830	246 GluGlyLeuGly-ProGlnValGluLeuProLeuTyrAsnGlnProSerAs 262	ò
US// CICGICGECGCCGCCCCCCCCCCCCCCCCCCCCCCCCCC	ACGGCACCGGCACCTCGGCGACCGATCGAGGCCCAGGCGCTCCTGGCGACTACG 77885	: 음
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:: 78650 GCCCTCTACCGCCTGGTCGCGTCCCTCGGC	186 ValGluGlnGlnIleSerLysLeuLysLysGlnGlnGlnGlnLeuGluGluGluAlaAla 205	ò
492 ysSer	77713 CGG 77736 Q	q
78590	GlulleThrMet 185	ò
	77653 CGCCCAGCGCGAGGGCCCCGGGTCCTCGCCGTGCTGCGGGCTCCGCCATCAACCAGGA 77712 Ov	g
	146 GluproValSerProProProHisThrAspProGluLeuGluLeuValProPro 165	ò
	77593 GGCCGACGGCACCGGCTGGGGGGCGCGGCATGCTGGTCCTGATGCGGCTCTCCGA 77652	ą
::: 	130 AlaGlySerGluAspLeuThrLysAspArgSerLeuThrGlyLysLeu 145	ò
	77560	qq
78414 GAACCGCCGAAAGGCCCTGCGCCGTCC	110 LeuGlüLeüLeüProAspProLeüLeüArgProSerProLeüLeüAlaThrGlyGlnPro 129	ò

ATGCCCAGGATCCCTGCCGGACGGCGCCGC CCCGAGGAGCTCTCCGCCGTCACCGGCCTGGC 79350 CGCTGGGCCGCCTCTTCGACGGCACCGGCGC 79410 :::||| -----TGGCGCTCATGGACCCGATG 79030 GCATCCCGGTCCTCCCAACCTCACCGGCACC 79111 |||::: | | || |::: | | | | |::: :::: ||||:::||||||| ccggggtctrctctcaaagacgccrgcacc 78769 AGGCCCTGCCGCGCGGCGCGATGCTCGCG 78829 |||||| ccrccgrcgrcgccggcaccaaggaagcc 78934 cccccagacccaagaccacccccccg 78991 ------GAATTCCGCGCCGTC 79051 SCGTCACCCCCGACTTCGTCGCGGCCACTCC 78709 cretrereegecagecreecagecreege 78473 ccegrerrceccececriceacaaaica 78533 snTrpSerAlaIleAlaArgMetValGly--- 642 AlaAlaSerGluGlu------Al 694 GluMetGluAlaSerGlyValSerGlyAsnGl 714 snPheryrPheAsnTyrLysLysArgGlnAsn 661 uGluLysProGluValGluAsnAspLysGlu 543 nrserGlyGluAspAsnAspGluLysGluAja 563 snSerGlnGlyArgArgLysGlyArgIleThr 583 luGluAlaIleThrProGlnGlnSerAlaGlu 603 erArgTrpThrGluGluGluMetGluThrAla 623 nglnglnproMetProArg----- 514 ----SerSerGlnGluGluLysAspGluLysG 524 MetGlnHisProLysAsnPheGlyLeuileA 455 ThrValAlaGluCysValLeuTyrTyrTyrL 472 LeuvalArgArgSerTyrArgArgArgGlyL 492 ------GlnGlnGlnGlnGlnG 499

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ALPHARMA AS.
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25351 GGACTGGACCCCGGTTCCCGACACCGCCAACTCCGCGCCGCCGCCGTCGG 25410
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                         2436 GluGlyAsp------CysAsnArgArgThrProLeuThr
                                                          2447 AsnArgValTrpGluAspArgProSerSerAlaGlySerThrProPheProTyrAsnPro
                                                                                            2467 Leu--------IleMetArgLeuGlnAlaGlyValMetAlaSerProProPro
                                                                                                            25519 Gerccccrcegceccccacrcsaccecacaccececaccacceccccccacce
                                                                                                                              -ProGlyLeuProAlaGlySerGlyProLeuAlaGlyProHisHisAla
                                        25411 CCCGGACACCGCCGTGCTCGCCGACGCCCTCGGCGACCCGGCCGTCGCACGCCACGCAAC
                                                                          ------caaccroscossacaccacccccccccccccccrcr
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/product= "NysD2 complete protein"
120628. .121308
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3492. .51099
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57503. .58687
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complement (58786. .58980)
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product= "NysJ protein"
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6337. .34771
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            The present invention relates to the cloning and sequencing of the gene cluster encoding a modular type I polyketide synthase (FKS) enzyme involved in the biosynthesis of the macrolide antibiotic nystatin. The nystatin PKS is useful as antifungal antibiotics. The present sequence is a Streptomyces noursei nystatin PKS gene cluster DNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  New nystatin polyketide synthase polynucleotides and polypeptides, useful as antibiotics and antifungals.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             -----LeuProGluLeuGlyLysSerGluMetGluPhelleGluSerLysArgProArg 109
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                                                                                                                                                                                                                                                                                                                                                                                                     WPI; 2001-557614/62.
P-PSDB; AAE10143, AAE10144, AAE10145, AAE10146, AAE10147, AAE10148.
AAE10149, AAE10150.
                                                                                                                                                                                                                                                                                                                                          Strom AR;
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Sletta H, Gulliksen O;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Claim 1; Page 188-254; 266pp; English.
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10-APR-2000; 2000GB-00008786.
14-APR-2000; 2000GB-00009387.
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29.72%
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DZIECLEWSKA H.
ZOTCHEV S B.
SEKUROVA O N.
FJAERVIK E.
STROM A R.
VALLA S.
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2169 AlaSerC 24316 CGCCTC 2179 Pro	24376 CCTCGAC	24436 CGCCGCC	2186		24556 CCTCTCC	2197 SerProH	24616 CCCGGGC 2217 GlyGlyG	: 24675 ACCTCGA	2237 SerArgS		24774 GCACCGI	2277 GluSer	24834	2297 HisAsn	24885 CGGTCTC 2317	24944 ACCACC	2323 GlyLeu	25004 GGCGAC	2342	25064 TTCGGC	25124 CGCGAC	2360	25184 GCGCTA	2378 ProAla		2398 GlyGly 	25252 C1-GGA 2418 Ala
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24496 C	CCCCTCCTCGGCGCCGCCGTGCAACTCCCCGACGACGGCGCACTCTTCACCGGCCG 24555	
2187	2196	
24556 0	2216	
16	CCCGGGCACCGCACTGGCGTCGCCGCGGGGGGAGA-CCGGCAGGGGCC 24674	
2217 0	GlyGlyGluAspGlyIleGluProValSerProProGluGlyMetThrGluProGlyHis 2236	
237	SerargseralaValTyrProLeuLeuTyrArgAspGlyGluGlnThrGluProSerArg 2256	73
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235	GlyLysTyrAspGlnTrpGluGlu	
12	GCGACACCGGACTCCCGGTTCTCCTGGGAAGGCGTCACICIGGCACGCCTCCGGGGGGGGGG	7
2360	524	143
2378	ProAlaAlaMetProIleThrAlaAlaAspGlyArgSerAspHisThrLeuThrSerPro 2397	97
2398	erSerArgLysAlaLyBSerPro 2417 :: ACCCGCGACGCCTCTTCCACCT 2535	
241	AlaProGlyLeuAlaSerGlyAspArgProProSerValSerSerValHisSer 2435	

1116 6	1136 1	20521 0	1156	20581	1175	20641 (1187	20701	1205	20761	1223	20820	1238	20880	1258	20912	1278	20943	1298	21003	1317	21030	1337	21063	1354	21109	1374	21144	1394	21189	1414	21244	1433	21276	1449
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	1175 -	ArgGlyGlnAlaGlyProProGluSerLeuGlyVal 1186
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18569 ATCCGCGCCACCGAGGACGAGGTCACCCCCCACCTCACCGACGACGTCTCG 18619 544 AspLeuLeuLysGluLysThrAspAspThrSerGlyGluAspAsnAspGluLysGluAla 563 ::: 18620ATCGCCGTCAACGGGCCCACCTCCGTCGTCGTCGTCGCCGCACCGAGGAAGCC 18673 564 ValAlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgLysGlyArglleThr 583	18674 Grédecagearesdesesecerreacesecédearescecedesereces 18730 584 ArgSerMetalaAenGlualaAenSerGluGluAlalleThrProGlnGlnSerAlaGlu 603 18731 GredeceAcecerecece	Ala GTC	Lystysciytecheduthisciyargashirpseralailealargaevaloiy bystysciytecheduthisciyargashirpseralitecheargaecvachili	643SerLysThrValSerGlnCyeLysAsnPheTyrPheAsnTyrLysLysArgGlnAsn 661	uargasnalaargarg ::: rgaccacgcrcgrcga	682 LysLysLysLysAla-ProAlaAlaAlaSerGluGluAl 694 ::: ::::: 1 18970 ACTCGCCCCGACGCCGTGCTGCCCAGGAATCCCTGCCGGACGCCCCC 19029	694 aAlaPheProProValValGluAepGluGluMetGluAlaSerGlyValSerGlyAenGl 714	714 uGluGluMetVal 718	718 718 718 718 718 718 718 718 718	719 -GluGluAlaGluAlaLeuHisAlaSerGlyAsnGluVal-ProArgGlyGluCysSer- 737	738	750 erileproSerProHisThrGlualaAlaLysAspThrGly763	764	776 ABPGLyProProPro	781GlyProProThrProProArgArgThrSerArgAlabrolleGluP 796	796 roThrProAlaSerGluAlaThrGlyAlaProThrProPro 809
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involved in the biosynthesis of the macrolide antibiotic nystatin. nystatin PKS is useful as antifungal antibiotics. The present seque a Streptomyces noursei nysl DNA of nystatin PKS gene cluster
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P-PSDB; AAE10125, AAE10126, AAE10127, AAE10128, AAE10129, AAE10130,
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2328 ArgSerGlnAlaValGlnGluHisAlaSerThrAsn	2378 ProAlaAllaMerProIleThrAlaAlaAspGlyAygSerAspHisThrLeuThrSerPro 2378 ProAlaAllaMerProIleThrAlaAlaAspGlyAygSerAspHisThrLeuThrSerPro 1 9052 AccGGTCTCGCCTGCGGCGAGCGCCCACAGGTCCTCCGGAGAGGTCACGCCACCC 2398 Gly	Qy 2418	Qy 2461 OPheProTyrAsnProLeuIleMetArgLeuGlnAlaGlyValMetAlaSerProProPr 2481 bb 8719 TCGGCCCGGGAATGTCCGAGCAGCAGCCG	RESULT 47 AAD17184 ID AAD17184 standard; DNA; 65140 BP. XX AC AAD17184; XX DT 29-NOV-2001 (first entry) XX XX DE Streptomyces noursei nysl DNA of nystatin PKS gene cluster. XX XX XX XX XX XX XX XX XX XX XX XX XX	20 = 1 = 1 = 1 = 1 = 1 = 1 = 1 = 1 = 1 =
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qq	14412 GACGCCCACTGGTACGACCCTGCCCATACGTCGCCAGCAACGCCTGAGCCTCGATCGGAT 14353	è	**************************************
ò	989 AlaAlaProThrLysProAlaProProAlaProProProProGlnAsnLeuGlnProGlu 1008	े र	
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QC .		ò	1493
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g (GGAAAGUACCCICACGCAIGIAIGACGIACCGGGCCCGGGICCGGAAICGGACCACAGGC	λō	1504 TyrGluG
S 6	12/2 INDVBLYBEGIVATIBVALLEGUSETIYTGLUGLYGIYMRCSETVALINICATNCYBSGELYBG 1292	q	12482 GTCGAGG
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WPI; 2000-365602/31. P-PSDB; AAY92707, AAY92708, AAY92709.

Recombinant DNA compound encoding oleandolide polyketide synthase for synthesizing polyketides comprising a coding sequence for a domain of loading module or any one of extender modules.

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Disclosure; Page 14-26; 86pp; English

The sign part of the Streptococcus antiboritus oleandowich gene cluster. The oleandolide polyketide synthase (PKS), also known as 8,8adeesignated oleah, oleahI and oleahII. The PKS is a type I "modular"
clesignated oleah, oleahI and oleahII. The PKS is a type I "modular"
clesignated oleah, oleahI and oleahII. The PKS is a type I "modular"
clenzyme, where each ORF encodes 2 term modules and the first ORF also
encodes the loading module. Each module is composed of at least a
cercynthase (KS), acyl-transferase (AT) and an acyl carrier protein
(ACP) domain. The oleandolide PKS loading module contains an inactivated
KS, called KS-Q, where Q is the abbreviation for glutamine, present
instead of the active site cysteine required for activity. The large
multifunctional PKS enzymes catalyze the biosynthesis of polyketide
macrolactores through multistep pathways involving decarboxylative
carbon processing activities. The macrolide product of the PKS, 8,8acarbon processing activities. The macrolide product of the PKS, 8,8acarbon processing activities. The macrolide product of the PKS, 8,8acarbon processing activities. The macrolide product of the PKS, 8,8acarbon processing activities. The macrolide product of the PKS, 8,8acarbon processing activities. The macrolide rounding an oleandolide, is further modilied by epoxidation and glycosylation of including an oleandolide PKS operably linked to a promoter. Also
including an oleandolide PKS operably linked to a promoter. Also
cincluding an oleandolide PKS operably linked to a promoter. Also
incapable of binding substrate (the KSI-o mutation). The particular, the
cincapable of binding substrate (the KSI-o mutation).
cin the codon for the active site cysteine. The oleandolide PKS is useful
continues. Heterologous expression of oleandolide pks in host cells such
as Streptomyces coelicolor and S. lividans is also made possible.
Communiar by Active and converted to the content of provided to cultures of
configurations of provided to culture and converted provided to cu of the Streptococcus antibioticus oleandomycin gene cluster derivatives of erythromycins A-D. (Updated on 06-AUG-2003 to correct OS

Sequence 50937 BP; 6672 A; 16253 C; 19272 G; 8740 T; 0 U; 0 Other;

CCCCCCGACGGACAACTC-----CGCGCCGCACCACCCACGACGCCATCCCCTCAC 17410 ArgAlaThrGluProArgTyrProProHisSerLeuSerTyrProValGlnIleAlaArg 32 sheuserProGlySerIleIleGlnProGlnArg-----ArgArgProSerLeuLe sSerTyrLeuProGluLeuGlyLysSerGluMetGluPhelleGluSerLysArgProAr ThrHisThrAspValGlyLeuLeuGluTyrGlnHisHisSerArgAsp-TyrAlaSerHi uSerGluPheGlnProGlyAsnGluArgSerGlnGluLeuHisLeuArgProGluSerHi caacgaacecrcaccacgacccacgcarcagccarcaccac 50937 684 286 1254 1041 Length: Matches: Conservative: Mismatches: Indels: (1-50937)Gaps: US-09-522-753-5 (1-2517) x AAA09469 7.03e-11 648.50 29.82% 21.03% 17352 C-----Percent Similarity: Best Local Similarity: Alignment Scores: Pred. No.: 13 33 25 17463 8 17490 69 17409 Query Match DB: Score: 요 8 셤 요 ઠે ò ò 셤

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16791 GATCCTCAAAACCAGCCGACGCCACAACACCCCACGAACCGCCGGCACACCTCCTCATCAA 16732 16731 CCCCCACCGGCAGGTCCACCAGCCCACCCACCCACTGGGGATGCTCCAGACCCAGAACAC 16672 CCCCGCTCACACCACGCCCCGCGACAGCCTCCGCATAAGCCCCGGCGTCAGGACGGGTCG 16390 -----CT 16366 17115 CCGGCTCACCCAGATCCGACAACAA......--CGCCCGCAACGCCACGAT 17071 CAGCCACATCACAGGCCCGCACCGACACCCCGAGCCCCAACCCCTCCAGCTCCCGCA 17011 16890 CCAACACCGTCCCACGCGG-----CCGCCAACCACCCCCACCACCATCCACCACAGAG 16837 16836 GCACCAGACGACGCACCCACAC. -----TCACCGCACCCGCCACCTTGGCCGCCATCACTCAGCCGCCTCCTGGACA 383 201 uGluGluAlaAlaLysPro-----ProGluProGluLysProValSer---ProPr 217 240 aAlaHisArgIleLeuGluGlyLeuGlyProGlnValGluLeuProLeuTyrAsnGlnPr 260 306 -MetGluAlaLeuGluLysLysValGluArgIleGluAsnAsnProArgArgArgAlaLy 325 gleuGluLeuProAspProLeuLeuArgProSerProLeuLeuAlaThrGly----17319 CATACACCGCCTGCCCCCC-----ACTGCCCCACACACCGGCATTGGAGGAGA ------GlnProAlaGlySerGluAspLeuThrLysAspArgSerLeuTh 17271 ACAACACAAACGCCTCCAGACCACAGGATCCACCAACTCACCCA-GAT----uLeuValProProArgLeuSerLysGluGluLeuIleGlnAsnMetAspArgValAspAr --AspGluAsnArgLysLysAlaGluAl 16950 CCCCACCACCACCAACCACCAGGCCGTATGCGCACCCAAACCACAAAGACCACAGACCACGTGA ulleteuTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPheCysGl 325 sGluSerLysValArgGluTyrTyrGluLysGlnPheProGluIleArgLysGlnArgGl 345 uLeuGlnGluArgMetGln----SerArgValGlyGlnArgGlySerGlyLeuSerMe ----- LeuTyrAspAlaAspGlnGlnArgIleLysPheIleAsnMetAsnGlyLeuMe gGlulleThrMetValGluGlnGlnIleSerLysLeuLysLysLysLysGlnGlnGlnLeuGl oSerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLysLe tSerAlaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSerGluGlnGl rGlyLysLeuGluProValSerProProSerProPro---HisThrAspProGluLeuGl 217 oProlleGluSerLysHisArgSerLeuValGlnIleIleTyr------uAsnLeuGluLysGlnMetArgGlnLeuAlaValIleProProMet----|||-:::||| |-:::||| GATCGAGCTCCACGACACTGAC nArgTyrAspGlnLeu----16491 GCTC----109 128 142 161 232 300 363 383 17070 260 16449

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oleandomycin; oleandolide; polyketide synthase; oleAI; oleAII; pKS; type I; 8,8a-deoxyoleandolide synthase; modular; ketosynthase; acyl-transferase; acyl carrier protein; inactivated; polyketide; macrolactone; antibiotic; motilide; erythromycin; ss.
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754	39 ProAlaThrValAsnSerSerAspThrGluSerIle-ProSerPro	7
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738	19 GluGluAlaGluAlaLeuHisAlaSerGlyAsnGluValProArgGlyGluCysSerGly	7
8946	63 CCCACGCCGTCCCGTCCG	89
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638	19 GluMetGluThrAlaLysLysGlyLeuLeuGluHisGlyArgAsnTrpSerAlaileAla	φ
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618	99 GlnGlnSerAlaGluLeuAlaSerMetGluLeuAsnGluSerSerArgTrpThrGluGlu	Ŋ
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           multi-functional proteins which direct the synthesis of the polyketide platenolide. Platenolide is the basic building block of the macrolide antibiotic spiramycin. The tyld gene (see AAT80413) is the tyldscone antibiotic spiramycin. The tyld gene (see AAT80413) is the tyldscone synthase gene cluster of the invention. The tyld sequence was isolated from Streptomyces fradiae, and encodes multifunctional proteins which direct the synthesis of the polyketide tyldscone. Tylactone is the basic building block of the antibiotic tylosin. The hybrid sequence can be used to transform S. ambofaciens lacking the srmG ORF1 sequence, or S. fradiae lacking the tylG ORF1 sequence, so that they can produce polyketides. The DNA sequence can be modified so as to alter the type of carboxylic acids incorporated, the number of carboxylic acids incorporated and/or the post condensation reactions performed, thereby resulting in novel tylosin-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          84 LeuArgProGluSerHisSerTyrLeuProGluLeuGlyLysSerGluMetGluPhelle 103
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ð i	GluGinHisHisileArgGlySerileThrGlnGly1leProArgSerTyrValGluAla 1366
a 8	1382
; A	CGGAGAGGGC
δ'n	1402
Ωp	3391CCATCAGCGCCGGATGCACGCCGTAG-TCCTCGGCGCTGTCCCCCGGGCCTTG 3339
λΌ	1403 GlyProLeuLysLeuLysProAlaHisGluGlyLeuValAlaThrValLysGluAlaGly 1422 Db
qq	3338 GGCAACACCACG 3327 QV
ò	1423 ArgSerIleHisGluIleProArgGluGluLeuArgHisThrProGluLeuProLeuAla 1442
qq	3326 cegecgaacaacercerecereceredaceceacacerceace-ceceregaa 3277
ò	
Db	
δý	pThrGlyAlaSerThrThrGlySerLysLysHisAspValArgSerLeulleGlySerPr
QQ	3216 CTCCTCCGCGCGCGCACTGGCCACGTCGCGAGCTCATCGAGCTCACC 3169

1776 1785 1725 jaAlaGlyProArgGlyIlelleAspLeuSerGlnValProHisLeuProValLeuValP 1745 1754 laMetAspArgLeuAlaTyrLeuProThrAlaPro-----GlnProPheSerSerArgH 1772 :::::||||||| 2355 CAGCTGCGTCGCCTTCGGCCTGAGCGACGACTTCGCTCCTAGCTCGGGCAGCAGCGC 2296 2517 CACCGGGTGCGCGC------TCACCTCCACGAACACACGTGCCCGTC 2476 roProThr-----ThrA 1754 gracregedercededecregededecacadetercerercadetreaderaaaacge 2356 2639 1515 1535 ysproArgGlnSerProLeuThrTyrGluAspHisGlyAlaProPheAlaGlyHisLeuP 1555 2941 roArgGlySerProValThrMetArgGluProThrProArgLeuGln-----G 1571 1571 luGlySerLeuSerSerLysAlaSerGlnAspArgLysLeuThrSerThrProArgG 1591 2831 1591 lulleAlaLysSerProHisSerThrValProGluHisHisProHisProlleSerProT 1611 ------chregeeereagechagetregtreechedeagearg 2788 1631 1631 laPheAspProThrSerIleProArgGlyIleProLeuAspAlaAlaAlaAlaTyrL 1651 1651 euprodrgHisLeudlaProdenProThrTyrProHisLeuTyrProProProTyrLeulleA 1671 1671 rgGlyTyrProAspThrAlaAlaLeuGluAsnArgGlnThrIleIleAsnAspTyrIleT 1691 1691 hrSerGlnGlnMetHisHisAsnThralaThrAlaMetAlaGlnArgAlaAspMetLeuA 1711 1498 3049 1515 jaSerSerGlyGlySerlleAjaArgGlyAlaProVallleValProGluLeuGlyL 1535 1611 yrGluHisLeuLeuArgGlyValSerGlyValAspLeuTyrArgSerHislleProLeuA ------checacerecaecececaeceretack--------------LeuSerProGlyGlyProThrHisLeuogly-----Arg-ThrPheProProValHisProLeuAspValMetAlaAspAlaArgA 2991 GGCCAAGACCATCTCCGACAGGGCCCCGGC-----TGCCCACGTGCGCCCAGCCGC 1498 laieuGluargala------ÇyeTyrGluGluSerLeuLysSerArgProGlyThrA 3108 TICHGGCCCCTGCTCCAGCTGTACAGCCCAAACTCACGCCGGCCCGCCGCGTCCGG 1772 isSerSerPro--------2295 1745 2692 2577 2415 2638 2605 1777 3048 1555 1480

Page 150

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more genes in the tmbA gene
 one or
enzymes comprises all or a portion of cluster of Sorangium cellulosum.
  4448400000000000
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Disclosure; SEQ ID NO 17; 73pp; English

The invention describes a recombinant DNA vector that comprises all or portion of one or more genes in the tubA gene cluster. Also disclosed in the host cell that comprises the tubA genes of Sorangium cellulosum or fragments of those genes. The DNA vector is useful in preparing hybrid polyketide synthase (PKS) enzymes and the polyketides produced by such hybrid enzymes. The gene products of the tubA cluster can be used to synthesize the polyketide combamycin. This sequence represents the Sorangium cellulosum tubA gene cluster tmbC open reading frame.

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Sequence 11358 BP; 1488 A; 3388 C; 4691 G; 1791 T; 0 U; 0 Other;

11358 659 271 1075 926 139		17 ProArgTyrProProHisSerLeuSerTyrProValGlnIleAlaArgThrHisThrAsp	GCGCAGCGCCACTACCCG	AspTyr	7341 CGCTCCCTCGCAAGCGACAAGCGCTCCGCACACCACCCCGCCGACACCTCGCCTGGCT
1139 659 271 1079 926 139		ılle	-	rArg	- - - - - - - - - - - - - - - - - - -
Length: Matches: Conservative: Mismatches: Indels: Gaps:	(1-11358)	LeuSerTyrProValGl1		HisHisSerArgAspTyr	cgcrccgcacacacaccg
1.73e-11 652.00 31.86% 22.58% 4.93%	US-09-522-753-5 (1-2517) x ADC26983 (1-11358)	rProProHisSer	7386 CCCCGACTGCTGCCGCACCGCCTGGCT	37 ValGlyLeuLeuGluTyrGln	ccccaaccacaca
Alignment Scores: Pred. No.: Score: Percent Similarity: Best Local Similarity: Query Match:	-753-5 (1-251	17 ProArgTy	7386 CCCCGACT	37 ValGlyLe	7341 CGCTCCCT
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7176 CGACGGCTGCTCCTCCCCCCACCACCGCACCGACCAC---AGACCAGCCCGT 7120 -- 6865 6821 G 7342 7119 ccacedececaececercarcecacecióca-ccécerccecaacecrececerecer 7061 ---HisThrAspPro---GluLeuGluLeuValProPro-ArgLeuSerLysGluGluLe 172 SerHisSerTyrLeuProGluLeuGlyLysSerGluMetGluPheIleGluSerLysArg 107 -----LeuleuArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySerGluAsp 134 LeuThrLysAspArgSerLeuThrGlyLysLeuGluProValSerProProPro 154 ulleGlnAsnMetAspArgValAspArgGluIleThrMetValGluGlnGlnIleSerLy 192 36 qr - 49 50 ---AlaSerHisLeuSerProGlySerIleIleGlnProGlnArgArgArgProSerLeu 68 69 LeuSerGluPheGlnProGlyAsnGluArgSerGlnGluLeuHisLeuArgPro---Glu 87 geneecceceákicicheccacaceccccacacacichicacacicichicascacacacacacaca sLeuLysLysLysGlnGlnGlnLeuGluGluGluAlaAlaLysProProGlu------ProGluLysProValSerProProFioIleGluSerLysHisArgSerLeuValGlnIl 5901 ccrgcaccgargcgcgaga-----ggcgaagrgcgrccgrg------ProArg-----LeuGluLeuLeuProAspPro----108 117 172 192 88 135 7012 155 6864 210 6961

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229	elleTyrAspGluAsnArgLysLysAlaGluAlaAlaHisArgIleLeuGluGluGl249 ::::::::::::::::::::::::::::::::::::
249	yProGlnValGluLeuProLeuTyrAenGlnProSerAspThrAr 264
264	gglnTyrHis
268	GluAsnIleLysIleAsnGlnAlaMetArgLysLysLeulleLeuTyrPheLy 285 ::: ::: ::: ::: ::: ::: :::
285	SARGARGASOHHISALAARGLYSGINTrpLySGINLYSPheCYSGINARGTYrASpGINLE 305
305	uMetGluAlaLeuGluLysLysValGluArgIleGluAsnAsnProArgArgArgAlaLy 325
325	sgluserLysValArggluTyrTyrGluLysGlnPheProGluleArgLysGlnArgGl 345 :::
345	uLeuGlnGluArgMetGlnSerArgValGlyGlnArgGlySerGlyLeuSerMetSerAl 365 ::::: ::: ::: ::: ::: ::: ::: ::: :::
365	aAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSerGluGlnGluAsnLe 385
385	ugluLysGlnMetArgGlnLeualaVallleProProMetLeuTyrAspAlaAspGlnGl 405
405	narglleLysPhelleAsnMetAsnGlyLeuMetAlaAspProMetLysValTyrLysAs 425
425	pArgGlnValMetAsnMetTrpSerGluGlnGluLysGluThrPheArgGluLysPheMe 445
445	tGlnHisProLysAsnPheGlyLeulleAlaSerPheLeuGluArgLysThrValAlaGl 465
465	uCysValLeuTyrTyrTyrLeuThrLysLysAsnGluAsnTyrLysSerLeuValArgAr 485 :: gaccGrccrGGaTCACCGCAGAGCCACGATCACCCCCAGCACACGGTCGCCGTC 6163
485	
505	InGlnGlnGlnGlnBroMetProArgSerSerGlnGluGluLysAspGluLysGluL 525
525	ysGluLysGluAlaGluLysGluGluGluLysPro536
537	GluValGluAsnaspLysGluAspLeuLeuLysGluLysThrAspAspThrSerGlyG 556

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1774Thr 1783		17277 GCGACTGGTCGACCACCGTCATGGACGCCGGCTACT		17436 ACATGATCGACGACGCGGAGTGGCGGTGTCGCCTCCGGCACCCTGCGCCGCGCGACAACG 17495 1883 MetLysGly1leIleThrAlaValGluProSerLysProThrValLeuArgSer 1900 1883 MetLysGly1leIleThrAlaValGluProSerLysProThrValLeuArgSer 1900 11496 GCGGTCTCGCTGCTGCTGTCCGCCGCGCGGGTGTTCGTCCGCGGTGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCCAGGTCGTCGTCCAGGTCGTCGTCCAGGTCGTCGTCCAGGTCGTCGTCCAGGTCGTCGTCCAGGTCGTCGTCCAGGTCGTCGTCCAGGTCGTCGTCAGGTCGTCGTCGAGGTCGTCGTCGTCGTCGTCGTCGTCGTCGTCGTCGTCGTC		varitricin bearcostations:	17661 TCACCGCGCCCCACCCCCCGCGT	1974 SerProSerLysGlySerGluProArgProLeuValProProValSerGlyHis 1991	 GCAAGAGGGCTCCACCGTCGACTCCTGGCGCTACCGCCCGACCTGGAAGCCCGTCACCA 	17862 AGCTCCCGCAGCGCACCCTGGACGGCTCTGGTGTCCGCCGACGGCGTCGACG 17921 2026 ThrGlnSerLysProPheSerlleGlnGluLeuGluLeuArgSerLeuGlyTyrHisGly 2045 2026 ThrGlnSerLysProPheSerlleGlnGluLeuGluLeuArgSerLeuGlyTyrHisGly 2045 2026 ThrGlnSerLysProPheSerlleGlnGluLeuGluLeuArgSerLeuGlyTyrHisGly 2045 17922 ACACCGACGCCCGGAGGCGCTGGAACGCGCGCGGGGCGCCGAAGGTGCGCCGGC 17972		18033 ACGGCCTGACCGGCATCGTCTCCGTCCTCGCGGCGCCCGAGCGGACCGGTGCCGTCCCCG 18092 2063 SerLeuThrHisAspLysGlyLeuProLysHisLeuGluGluLeuAspLysSer 2080
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1474	16238 GCCCACGTCATCCT 1505 luGluSerLeuLysSerArgProGlyThrAlaSerSerGlyGlySerIleA 1505 luGluSerCeuLysSerArgProGlyThrAlaSerSerGlyGlySerIleA 16277 GCCGAGGAGCAGCAGCCCGGGCGCCCCTGCCGGTCGCCGGGTGGCGTGGCGTGGCGTGGCGTGGCGTGGCGTGGCGTGGCGTGGCGTGGCGTGGCGGTGGCGTGGCGGTGGCGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGCGGTGGGGGG	1523 laargGlyAlaPrcValileValProGluLeuGlyLySPrAkrgGlnSerProL		1563 rgGlurrollir.Floatybedollir.commigned for the fill for the following for the fill for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the following for the followi	16566GTGTCGCGGACGTCGCGGGCCGCACCGTCTTCGTCTTCCCCGGCCA 16611 1603 isHisProHisProIleSerProTyrGluHisLeuLeuArgGlyValSerGlyValAspL 1623 16612 GGGCTCC	1623 euTyrargSerHisIleProLeuAlaPheAspProThrSerIleProArgGlyIleP	1642 16681 1662	b 16720 CTCATCGACGTCTGCGGGGGCGCCAAGGGCGCCCCGAAGGCGCGTCGACGT 16776 y 1682 ArgGluthrileileAenAspTyrileThrSerGluGinMetHisHisAsnThr 1699 b 16777 CGTCCAGCCTCGTTCGCCGTGATGGTCCCTGGCCGCGTGTGGCGGGCCCA 16833	r	1/20 Deuprovalleuvalpro		dy 1759 Alaiyrbeurfolmfalario Db 17064Acgecececececrecresses 1711
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Ор	12991 -GCCGCAGCTGCCCTCCGCCGCCCTCCCCGGCCGACCGGCCGACCAGAAGCTGT 13049	ò
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654	eAsnTyrLysLysArgGlnAsnLeuAspGluIleLeuGlnGlnHisLysLeuLysMetG 674
13243	AG 13
674	luLysGluArgAsnAlaArgArgLysLysLysLysAlaProAlaA 689 ::::
68	laalaSerGluGluAlaAlaPheProProValValGluAspGluGluMetGluAlaSerG
13354	CCTCGTCCTGGACGCCGACT
709	lyValSerGlyAsnGluGluGluMetValGluGluAlaGluAlaLeuHisAlaSerGlyA 729
13373	1.0
72	snGluvalproArgGlyGluCysSerGlyProAlaThrValAsnAsnSerSe
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749	INSerIleProSerProHisThrGluAlaAlaLySASpinrolyGluAlaBiglyFrouyse
76	
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808	9 roProAlaProProSerProSerAlaProProProValValProLysGluGluLysGluG 829
1354(13
828	9 luGluThralaAlaAlaProProValGluGluGlyGluGluGluGlnLysProProAlaAlaG 849
1360	O CGACCCGGTCACCCACCCGCCCAGGCCACCGCCTGGGG 13638
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1363	 catgegececedanetrescectedaacacececececetargegegegegecetegecetece 13698
98	880
1369	9 gecceartrearcecegegegecedaragedectregeceaagereregegereaagaege 13758
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1375	9 ccccgacggcggaccaggrcgcccrgcgdgccaccggrgrgcacggccgccgccrggr 13:
06	0 erGlyArgalaThrThrAlaLysSerSerGlyAlaProGlnAspSerA 916
1381	
91	6 spSerSerAlaThrCysSerAlaAspGluValAspGluAlaGluGlyGlyAspLysAsnA 936
1387	9 GATCACCGGCGCCACCGGCGGCCTCG
93	6 rgLeuLeuSerProArgProSerLeuLeuThrProThrGlyAspP 951
1390	5 GCGCCGAGACCGCCCGCTGGCTCGCCCGCCGCCCCCCCCC
95	1 roArgAlaAsnAlaSerProGlnLysProLeuAspLeuLysG 965
1395	7 caccadececededececeddaecececedececececedaacrecececedaacredaaga 14
6	
1401	7 GIVGGGCCCAAAGCGIVIVAAILOGGCCTGCCAACGTCCCCCGGCCCGGCCCGGCCGGCCGGCC
86	5 roArqGluAspAlaAlaProThrLysFroAlaFro

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The invention relates to the gene cluster encoding the polypeptides responsible for the biosynthesis of the polyene antibiotic amphotericin (amph) of Streptomyces nodosus. Polynucleotides of the invention are useful for preparing amphotericin derivatives or analogue antibiotic agents with altered properties and in the biosynthesis of polyketides other than amphotericin. amphDIII, amphDII or amphDI mutants are useful for producing amphotericin derivatives glycosylated with alternative sugars; amphDIII or amphDII or amphDIII or amphDIII or amphDIII or amphDIII or amphDIII and perosaminyl-16-descarboxyl-16-methyl amphoteronolide B; amphDIII or amphDII and amphDI gene sequences are useful in the engineered biosynthesis of perosaminyl-16-descarboxyl-16-methyl amphoteronolide B; amphDIII, amphDII and amphDI gene sequences are useful for preparing polypeptides capable of addition of mycosamine to a polyketide other than amphoteronolide A or
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    for in vitro synthesis of GDP-mycosamine.
                                                                                                                                                                                                                                                                                                                                                       amphB
                   encoded by S. nodosus amphN
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Novel cytochrome P450 enzyme and nucleotides encoding the enzyme, use: for preparing amphotericin derivative or analog antibiotic agent with altered properties, in biosynthesis of polyketide other than
                                                                                                                                                                    nodosus
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            P-PSDB; AAE36116, AAE36117, AAE36118, AAE36119, AAE36120, AAE36121,
AAE36122, AAE36123; AAE36124, AAE36125, AAE36126, AAE36127, AAE36128,
AAE36129, AAE36130, AAE36131, AAE36132.
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                                                                                                                                                                                                                                                                                                                                  /product= "Polyketide synthase multienzyme housing extension modules 1 and 2 encoded by S. nodosus by
                                                                                                                                                                                                                                                                                                                                                                                                                           /product= "Polyketide synthase multienzyme housing extension modules 3, 4, 5, 6, 7 and 8 encoded by S.
                                                                                                                                                                                                                                                       product= "Polyketide synthase multienzyme housing
loading module encoded by S. nodosus amphA gene"
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  B or for preparing polypeptides for in vitro synthesis of GDP-mycc
The present sequence is S. nodosus amph biosynthetic gene cluster
                                                                                        product= "NDP-sugar aminotransferase encoded by
                                                                                                                                                                "Glycosyl transferase encoded by S.
                                                                                                                                                                                               /transl_except= (pos:65773. .65775, aa:Met)
66081. 70319
                 products "Cytochrome P450
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complement (64324. .65775)
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BP; 14248 A; 45141 C; 38354 G; 15450 T; 0 U; 0 Other;

Sequence 113193

Alignment Scores:

Length: Matches: Conservative:

8.5e-11 654.50 30.30%

Percent Similarity:

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11581 CTCCTTCGGCATCAGCGGCACCAATGTCCACACCGTCCTGGAGCAGGCGCCCGGGGACAC 11640
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                                                                                                          ----AlaSerHis---
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		λŏ	1303 uThrAlaAlaProLysArgThrTyrAsp
	985	qq	39472 CATCGGCGCCGCAGCAGCACCTGCGT
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                                    ----CysProValLeuAspLeuArgArgProProS
                                                                       CGGATCACCCAGCCGCGTCCCCATGCGCCTCCACCACATCCACATCCGCCACGT
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                                    2164 yrSerPheProGlyAlaSer----
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1516 erSerSerGlyGlySerIle ::::::: 8016 CGGCCGCTCGGGGTCCATG 1527 roValIleValProGluLeu 7956 CCGCCGCCTCTTTTTTTTTTTTTTTTTTTTTTTTTTTT	Oy 1552	Db 7677 CGTCCAGAATCCCGGCGGTGTGGAAG Oy 1597HisSerThrValProGluHi Db 7617 CGAGCGCTTCGGCTCGGCCACGTCA Oy 1615 Leu Db 7560 CTCGCAGTTCCTCGGCGAGTTCGGCG Oy 1616	7500 GGACCAGGCGTT 1632 PheAspPro 1649 TYTTYLEUPTO 7384 GCCGCGC-CCG 1669 LeulleArgGly 1669 LeulleArgGly 7349CGAGGC	7343 1709 7301 1726 7242 1736 7182
8927 GGCGTACTCCTGATGGGTCAGCCCGAC	uSerTyrGluGlyGlyMetSerValThrGlnCyBSerLygGluAspGlyArgSer-SerS 1298	1342 8532 1362 8527 1382 8497	1402 8437 1422 8404 1438 8344 8284	1453 InGlyThrProLeuLysTyrAspThrGlyAlaSerThrThrGlySerLysLysHisAspV 1473 19283 AGGCGCCTCGGTACGG 1
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1516 8016	erSerSerGlyGlySerIleAlaArgGlyAlaP 1527	~ ~
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1547	yalaproPheAla	
1552		.0 .0
1565	rothrProArgLeuGlnGluGlySerLeuSerSerSerSerLysAlaSerGlnAspArgLysL 1585	10 10
1585 7735	euthrserthrProArgGluIleAlaLys-SerPro	~
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7677	CGTCCAGAATCCCGGCGGTGTGGAAGACGGCGTTCGGCGGATACGCGGTGACGAGCGCGG 7618	~
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1615		10 -
7560	CICGCAGTICCTCGGCGAGTICGGCGGCGCCCGGCCTCCGGTCCGCGACGGCTGGTGA 750	_
1616 7500	ArgGlyValSerGlyValAspLeuTyrArgSerHisIleProLeuAla 1631	_ ~
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1649	TyrTyrLeuProArgHisLeuAlaProAsnProThrTyrProHisLeuTyrProProTyr 166	m -
7384	GCCGCGGC	·
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7343		C)
1709	MetLeuArgGlyLeuSerProArgGluSerSerLeuAlaLeuAsnTyrAla 1725	10 m
1726	AlaGly	10 00
1736	GlnValProHisLeuProValLeuValProProThrProGlyThrProAlaThrAla 1754	***
1755	MetAspArgLeuAlaTyrLeuProThrAlaProGlnProPheSerSerArgHisSerSer 177	5 #
7122		m

q	12797 GGCGGTGCCGCCGCAGGGAGAGCTGGCCGGCGCCGTCGAGGACCGCGCG 12747	ě	E
ò	117 -LeuLeuArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySerGluAspLeuTh 136	ें ह	
q	12746 CAGCGCGAGGGCAGCGCGGTGCGTCGTCCGCGTCGTCGTCGTGTCGAGGGGGAA 12687	2 2	7 /1071
δ	136 ILys-AspArgSerLeuThrGlyLysLeuGlubroValSerProSerProBroHisT 156	중 <u>음</u>	11963 CACACC
qq	12686 GCGGCCCGGGTGCTCGGACCGCGGACCTCACCAGGCCCCACACCCCCGGACA 12633	ò	502
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; f	::::::::::::::::::::::::::::::::::::::	qa	11844 CCAGGG
8 8	vsGlnGlnGlnLvsProVroGluProGluProGluProGluProValSerP	&.	531 Lys
7 A		qa	11784 AGGCCC
ò	216 roProProlle-GluSerLysHisArgSerLeuValGlnIleIleTyrAspGluAsnArg 235	ð i	539
qq		දු දි	
ò	236 LysLysAlaGluAlaAlaHisArglleLeuGluGlyLeuGlyProGlnValGluLeuPro 255	Š É	SSS INTSEL
q a	12500	o ,	
ò	256 LeuTyrAsnGlnProSerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAla 275	8 8	573 Asnser
đ	12473CCCGGAGAGCACCGGCAGTACGACCGCTTCCGGCGGCGCGCCC 12429	3 8	1884
ờ	276 MetArgLysLysLysLeulleLeuTyrPheLysArgArgAshisAlaArgLysGlnTrpLys 295	\$ E	11544 CTTCAC
qq	12428 CGCCGCCAGCGCCTTCCGCAGCGCCCGCACCTCCGGGTACAGCGCGGT 12381	àà	
ď	296 GlnLysPheCysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGluArg 315	<u> </u>	11485 274000
ΩP	12380 GGGCCCACCGTCCGGCTGATCCCCCCAGGCGCGTCCCGGCGGAGCCGGTCCGCGTCGGG 12321	3 8	
ò	316 IleGluhenAsnProArgArgArgAlaLysGluSerLysValArgGlu 331	<u> </u>	
q	12320 GCCGAGCACCGCCCAGGAGGCGGGGGGGGGGGGGGGGG	3 8	1112 (60000
ò	332 TyrTyrGluLysGlnPheProGlulleArgLysGlnArgGluLeuGlnGluArgMetGln 351	Š	013 4 25 4 1 1 5 4 5 5 4 1 1 1 1 1 1 1 1 1 1 1
qq	12260 GCGGTACAGCGACCGTTCGCCGCGTCCGTGCCAGGTCCAGCTTCCGGCGCGC 12201	3 8	
ò	352 SerargValGlyGlnArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluHisGlu 371	ž 2	
q		3 8	
ò	372 ValSerGluIleIleAspGlyLeuSerGluGluGluAsnLeuGluLysGlnMetArgGln 391	3 6	
q	12161 CGCGCCGCACGTCTCTGCGCCTGCAG12135	3 8	ASTOCK CEST
ò	392 LeuAlaValIleProProMetLeuTyrAspAlaAspGlnGlnArgIleLysPheIleAsn 411	3 E	11185 SECTION 1118
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ò	412 MetAsnGlyLeuMetAlaAspProMetLysValTyrLysAspArgGlnValMetAsnMet 431	à 8	839 AIGMEC
qq	12134	à	659 AraGln
ò	432 TrpSerGluGlnGluLysGluThrPheArgGluLysPheMetGlnHisProLysAsnPhe 451	. 'Ę	
đ	12110 CGTCAGCCGCACCCGCAGAGCACCCGCACCCGTCGCGTAGAGCGACACTCC 12060	}	
ò	452 GlyLeulleAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyr 471	; E	
οg	12059	}	

472	LeuThrLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyrArgArg 489 ::::::
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502	510
11903	CGCGTCCGGCTGCCGCCATACGGCGCGCGCAAACGCCT-CACCGTACTCGTAGC 11845
511	ProMetProArgSerSerGinGluGluLysAspGluLysGluLysGluLysGluAlaGlu 530
r .	1 0
531 11784	LysGluGluGluLysProGluVal
539	GluAsnAspLysGluAspLeuLeuLysGluLysThrAspAsp 552
	572
11664	CTTCCGGGCCCAAGTGAATGGCGACCCTCCGCCGTCCCAACTCGTCGGCGGACAACCC 11605
573	AsnSerGlnGlyArgArgLyArgIleThr583
11604	
584	
596	GlnGlnSerAlaglu 603
11485	GTACCCGGCAGCAACGTCACCCCGGACACGGCATGGTCCGTCAGCCAGTCGTGCTCGGAG 11426
604	LeuAlaSerMetGluLeuAsn GluSerSerArg 614
11425	CCCGCCAGCCGCCCGGAACACCCGGTCACCGGAGTCGGCCAGGGGTAACCCGCCACGG 11366
615	11
11365	AGGAAGGGGTGGTCCTCCCAGGTCATACCGAACCGGCCCGCAGCCGCGTCGCCGACTCCG 11306
618	618
11305	GCGGGGGATGTCACGGCCCAGTAGTGGTGGTGGTTGAAGGGGTAGGTGGGGAGGGTGGT
618	618
11245	AGGTGACCTGGGGGCAGGGCGGGGTCCCAGGTGAGGGTGGCGCCGTGGGCCCCAGGCCTCG 11186
619	GlumetGluThrAlaLysLysGlyLeuLeuGluHisGlyArgAsnTrpSerAlaIleAla 638 :::
639	
N	TGGACACTCCGGCGCGCTCGGCCGTCTGCTC-AATGGCATGGACGAGTACGGGTG 11067
659	ArgGlnAsnLeuAspGluIleLeuGlnGln-HisLysLeuLysMetGl 674 :::
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674	LLYBGIDARGABIALIAARGARGARGARGABLYBALBARGARGARGARGARGABGARGABARGABARGABARG

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||||||| cGGGCCAGCAGCCGCCGAGCGTCGCGCCCGTCACCAGGACCGTGCCCGCGG 12858
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         This sequence represents the tylactone synthase gene cluster of the invention. This sequence is also referred to as the tylg gene, and was isolated from Streptomyces fradiae. This sequence encodes multifunctional proteins which direct the synthesis of the polyketide tylactone, isolated from Streptomyces fradiae. Tylactone is the basic building block of the antibiotic tylosin. The DNA sequence can be modified so as to alter the type of carboxylic acids incorporated, the number of carboxylic acids incorporated and/or the post-condensation reactions performed, thereby resulting in novel tylosin-related polyketides
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        82 LeuHisLeuArg-ProGluSerHisSerTyrLeuProGluLeuGlyLysSerGluMetGl 101
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      74 ProGly-----AsnGluArgSerGlnGlu
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Sequence 43280 BP; 5269 A; 15187 C; 16651 G; 6173 T; 0 U; 0 Other;
                                                                                                     ot= (pos: 20010. .20012, aa: Met)
encodes protein shown in AAW22603"
                                                                                                                                                        shown in AAW22604"
                                                                                                                                                                                            "ORFS encodes protein shown in AAW22605"
pt= (pos: 816. .818, aa: Met)
encodes protein shown in AAW22601"
                                         /*tag= b
/transl_except= (pos: 14351. .14353, aa: Met)
/note= "ORF2 encodes protein shown in AAW22602"
                                                                                                                                                                                                                                                                                                                                                                                                                   DNA encoding Streptomyces fradiae tylactone synthase domain production of tylosin-related polyketide compounds.
                                                                                                                                                                                                                                                                                                                                                                                         P-PSDB; AAW22601, AAW22602, AAW22603, AAW22604, AAW22605
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Matches:
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                                                        2293 sLeuAsnThrHisAsnArgAsnGluProGluTyrAsnIleSerGlnProGlyThrGluIl
                                                                                           ---LeuMetThrTyrAr
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                                                                                                                                            2328 gSerGlnAlaValGlnGluHisAlaSerThrAsnMetGlyLeuGluAlaIleIleArgLy
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                                                                                                                                                                                                                       55178 CTCAGCAACGGCACGCATCACCGCA---GÁGGACAĞTCCACCGGTGTCAACA-----
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                                                                                         2313 ePheAsnMetProAlalleThrGlyThrGly----
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1225 TytatgdlySerIllethr————HisolythrPoolabspvalLeutyLyadly 6929 TC-dOrceanCorchAccorchCACACACACCACCACCACCACCACCACCACCACCACCAC	4 6 8 6 8 6 8	8 8 8 8 8	8888	8 6 8 6 8 6 8	8 6 8 6 8 6	868686
		1277 ValLeuSerTyrGluGlyGlyMetSerVal-ThrGlnCysSerLysGluAspGlyArgSe :::	1336 OPTOGLUARGUARGGAATUGCACACACCCACCCGACCCGGATACACATCCC 1336 OPTOGLUARGHISSERPTOHISHISLEULYSCIUGINHISHIS 1351 OPTOGLUARGHISSERPTOHISHISLEULYSCIUGINHISHIS	68848	68715	68568 TCACCACCCACACCACACTCCGCCCCCACCAACCCACCAACCACCGCCCCCCAACCAA

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929 AlaGluGlyGlyAspLysAsharg	70
937	roThr 948
70256 TTCCTCAAGGAACCACAGACGCTGAGCGAACGACAACGGCTC	CGACCCGT
94	AsnAlaSerProGlnLysPro 960
70202 CCGCTCCCGTGCCACCAGAGCACCCGCACCCCAAACCCGGACCGCTCCC	Ť O
961 LeuAspLeuLysGlnLeuLysGlnArgAlaAlaAla1lePro 111	AlaalailePro 974 CACACCCGCTCAACACCCA 70084
975ProlleGlnValThrLysValHisGluProProArgGluAspAlaAlaProThrLys	993
70083 ACACCGCACGGACACGCGAAATCAGACGAGTCGCCAGAAGAGAATGACCA	00 0
994	ProproProProGlnAsnLeu 1005 CTCCGCGAACAGACCACACA 69964
1006	SerProArgGly 1021
69963 ACACCTCCTCCGGGGGGGGGGCGCCGGTAGCCCGTGCCCGCCTCCACCTCCGGCA	TGCCCGCCTCCACCTCCGGCA 69904
1022 LysSerArgSerProAlaProProAlaAspLysGluAlaPheAlaAlaGluAlaGluLys	PheAlaAlaGluAlaGlnLys 1041
69903 CCGGAAGCGCCCCCGTCCACCTTCCCGTTCGGCGTCAACGGGATCTCA	ACCGCATCTCATCCAGAACCA 69844
1042 LeuProGlyAspProProCysTrpThrSerGlyLeuProPhePro	PhePro 1056
1057ValProProAr	7
83 AACGAAGACCATCCCCACCAACACCCACCGCAGTCACGTA	 raagccaccaaccggrgarcac 69724
1064 LysAlaSerProHisAlaProAspProSerAl	ProSerAlaPheSerTyr 1077
1078 AlaproProGlyHisproLeuProLeuGlyLeuHisAspThrAla	oThrala 1092
69663 CACCCICCACCTGACCGAACACGGAAACCACGGACCTICACCTGATCAT	::: ACCTTCACCTGATCATCGCAC 69604
1093ArgProValLeuProArgProProThrileSerAsnProProPro	eSerAsnProProLeulle 1109
69603 GGCCAACGAACTCAAGCGCGCCGTCCACCGGCCACCGCACCAAATCCCCC	ACCAAATCCCCCGTGC 69550
ValLeuGluArgGlnIleGlyAlaIleS	GlyAlaI
69549 GATACATACGCTCACCCGGCA	
1130 SerValGlnLeuHisValProTyrSerGluHisAlaLysAlaProValGlyProValThr	1149
69528CACCGAACGGACACGCCACAAACCGCTCCGCCCACAG	rcceccercagacceacece 694
1150 MetGlyLeuProLeuProMetAspProLysLysLeuAlaProPheSerG	aProPheSerGlyValLysGln 1169
69480 CGAGATAGCCCCGCCCAGACCCA	694
luGlnL	1189
69456CACCGCCACATACAACTCACCACCACACGGCACGG	4
1190GlnGluAlaSerValLeuArgGlyThrAlaLeuGlySerValPro	1204
	693
	1ProSerAspSerAlaIleThr 1224

dB.

anti-cholesterolemic; agrochemical; gene;

Streptomyces roseosporus

WO200259322-A2

01-AUG-2002

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Percent Similarity:
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2225 ValSerProProGluGlyMetThrGluProGlyHisSerArgSerAlaValTyrProLeu 2244
                                             etProAlalleThrGlyThrGlyLeuMetThrTyrArgSerGlnAlaValGlnGluHisA
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ID ABQ78872 standard; DNA; 90600
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The invention relates to a novel isolated nucleic acid molecule comprising a sequence that encodes a thicesterase or thicesterase domain, derived from a bacterial daptomycin biosynthetic gene cluster. The proteins of the invention have antibacterial, fungicide, virucide, antiparasitic, immunomodulator, antilipemic, and cytostatic activity. The polynucleotides may have a use in gene therapy. The compositions and methods of the present invention are useful for generating novel linear and cyclic peptides and improving yield of a product in a cell expressing an daptomycin non-ribosomal peptide synthetase (NRFS) to be used as new compounds or in producing new compounds, such as antibiotics, antifungals, antivirals, antiparasitics, antimitotics, antitumour agents, immunomodulatory agents, anti-cholesterolemic agents, siderophores, of agrachmicals and cytostatics. The sequence represents the 90kb region of the S. roseosporus daptomycin biosynthetic gene cluster

90600 BP; 12671 A; 32312 C; 31571 G; 14046 T; 0 U; 0 Other;

Sequence

301 1199 1091 142

Length:
Matches:
Conservative:
Mismatches:
Indels:

08e-11 662.50 30.47% 21.29% 5.01%

.. No.:

Similarity:

Isolated nucleic acid molecule from a bacterial daptomycin biosynthetic gene cluster encoding a thioesterase or thioesterase domain, useful for generating novel linear and cyclic peptides, and products in a cell.

Claim 7; Page 142-165; 227pp; English.

Silva CJ;

Baltz RH,

BRIAN P. BALTZ R H. SILVA C J. MIAO V P W.

MIAO/) (BRIA/) (BALT/) SILV/)

WPI; 2002-599794/64. Miao VPW, Brian P,

17-OCT-2000; 2000US-0240879P. 28-FEB-2001; 2001US-0272207P. 06-AUG-2001; 2001US-0310385P.

17-OCT-2001; 2001WO-US032354.

73562 GAGGGTGCGGAGGCTTTCGTGGCGGGGATGACATCAATGAGCGCCTGGTGCAGCGCTTC 73503 73563 65 73 ::: |||||| ||||||| |||||| ::: ValGlnIleAlaArgThrHisThrAspValGlyLeuLeuGluTyrGlnHisHisSerArg GTTCAGTTTCGCGACGAGGGTGTCGGCGTCGTCTGTACGACGGGAAGAGCCGGTCGCC 73502 GACGTCCAGAACACCGGCCAGACGCACGCCATCGGAATGTTGTACGCAGCACCGGGCCC ----ProSerLeuLeuSerGluPheGln-48 AspTyrAlaSer-----HisLeuSerProGlySerllelle---GlnProGlnArgArgArg--US-09-522-753-5 (1-2517) x ABQ78872 (1-90600) 28 73682 29 음. 장 용 ਨੇ à 엄 ò 셤 ò

Daptomycin biosynthetic gene cluster; thioesterase; antibacterial; fundicide; virucide; antiparasitic; immunomodulator; antilipemic; cytostatic; gene therapy; antimitotic; immunomodulatory; siderophore;

roseosporus daptomycin biosynthetic gene cluster 90kb region.

(first entry)

23-OCT-2002

ABQ78872;

BP

Qy 1896 hrValleuArgSerThrSerf Db 4062 CGTTC Qy 1916 laThrHisCysProLeuGlyGlyThrI Db 4101 CGCCGCCGCCGCCGTCACCGACCACCGACCA	Oy 1936 alleuleuproLysGlud Db 4161 CGTTCGCGCCAGCCACGCTGGCGTCGC Qy 1950ArgProArgAlaAsp7 Db 4221 TGTGCACGGTAGGCGCGGTCGCGCCGG	1966 4280 . 1986 4326	2006 4380 2026 4421	Oy 2046 erSerTyrSerProGluGlyValGlul	Oy 2086 euArgProLysGlnProGlyProVall	0y 2134 hrLeuAlaGlnHisIleSer(Db 4710 CGGTCATATCGGCATAGCCGGCACCC Oy 2152 roGlnGlnLeuSerAlaProLeuPro CGTCACCGACCCACCCGCCGTT
CCGTTGGCGCCGCCCACCACCACCACCACCACACACACAC	1597 HisSerThrValProGluHisHisProHisProIleSerProTyrGluHisLeuLeuArcus 3378 1597 HisSerThrValProGluHisHisProHisProIleSerProTyrGluHisLeuLeuArg 1616 1377 TTGCCACCGTTACCACGCTGCCGCCGCTGCGGGCCCGTGG	Ilepro-ArgdlylleproLeuAspAlaAlaAlaAlaAryrTyrLeuProArgHisLeuAl CGGCCGAGGCCCGGCGCGGGGTCACCGCCGTCCCCCCCCC	1672 yTyrProAspThrAlaAlaLeuGluAsnArgGlnThrIleIleAsnAspTyrIleThrSe 1692		aThralametaspargLeualaTyrLeuproThralaproGlaproPheSerSerArgH cgrGcCGG	1812 y8SerIleLeuThrSerThrThrValGluHisAlaProlleTrpArgProGlyThrG
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	1896	hrvalLeuArgSerThrSerThrSerSerProValArgProAlaAlaThrPheProProA 1916
	1916	lathrHisCysProLeuGlyGlyThrLeuAspGlyValTyrProThrLeuMetGluProV 1936::
	1936	alLeuLeuProLysGluAlaProArgValAlaArgProGlu 1949
	1950	ArgProArgAlaAspThrGlyHisAlaPheLeuAlaLysProProAlaA 1966
	1966	rgSerGlyLeuGluProAlaSerSerProSerLysGlySerGluProArgProLeuValP 1986
•	1986 4326	ProValSerGlyHisAlaThrIleAlaArgThrProAlaLysAsnLe
	2006	<u>ي</u> :
	2026	yTyrHisGly
	2046	<pre>srTyrSerProGluGlyValGluProValSerProValSerSerFroSerLeuThr srTyrSerProGluGlyValGluProValSerProValSerSerCerThr </pre>
	2066	8AsplysGlyLeuProLysHisL ACCGCCGGCTCCGCCATCA
	2086	ValLysLeuGlyGlyGluAl CCGGC
	2106	euargProLeuProGluSerGlnProSer
	2116	-SerSerProLeuLeuGlnThrAl :::: GGCCCGGCGTTACCTCCATCCTC
	2134	hrLeuAlaGlnHisIleSerGluValIleThrGlnAspTyrThrArgHisHisP 2152
	2152	1 0
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	2185	eupro-ProProAspHisGlyAlaProAlaArgGlySerProHisSerGluGlyGlyLys 2204
	2205	ArgSerProGluProAsnLysThrSerValLeuGlyGlyGlyGluAspGlyIleGluPro 2224

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	LeuLysAlaGluLysLysGluGlyGly :::	අු	253.
	CCGCCGTCGCCTCCGTCACCGCCGGGGGCGCCGAAGCCGGCCTT	ò	122]
	SerGlyArgAlaThrThrAlaLysSerSerGlyAlaProGlnAspSer	qq	2597
П	GCCGTGCGCGCCACTTGTGGAACCGAAACCGCCTTGTCCCCCCGGGGCGGCCCCCACCCGC	ò	124
•	AspserSerAlaThrCysSerAlaAspGluValAspGluAlaGluGlyGlyAspLysAsm	අු	264
	GTCGCCGCCGTCACCTCGCCGCCAGGCTCTCCGTCAAAATCCGCAAGATAGGT	λ̈́ο	126
8 1	936 ArgLeuLeuSer-ProArgProSerLeuLeuThrProThrGlyAspProArgAlaAshAl 955 :::::::::::: ::: ::::::::::::	අු	268
-1	AANGCCGICACCAAGCCAACAAAAACCAACCAACCCGCCCGGCCGG	ò	128
·	955 aSerFroGinLysFroLeuAspleuLysGinLeuLysGinArghiaAlatieFroFr 975	q	274
	GICCCGCCGAACGCCT	ò	130
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	LeuGinProGluSerAspAlaProGinGinFroGiySerSerFroArgGiyDy	QQ	288
	AHCGCCG11ACCGCCGACCCCACCACCACCACCATCCGCCTTCCGGCCCTTCCGGCCGTCGTCGCCGTCGACCATCGCCCGTCGATCGA	ολ	136
	BSETATG	qa	291
	GICGCCGGICTCCGCCGGCCGCCACCCGCCCCGTCGCCGTCGCCTCCCGCCCG	ð	138
	1031ABPLYBGLUALARNAAATAGLUALAGUTILIKSLEURFOGIJASPRI 1040	Ωp	297
	GCCGCCGTCGAMGCCCGTCGAMGCCCTTTAGGTCGGGGCGACCGACCAACCCGGGCCGACCAACCA	δλ	140
	O	qq	303
	GGCCGCCGCTTGCCTTCCCTGGCCGG	ò	142
i i	106Z 111ELYBALGSEIZIOTESAGAFIONESFICOSEIAGAFIGGGGIIJY-DIGHT-CEIOGI-7-11 - 100Z	qq	304
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		ò	148
	leGlyAlaIleSerGlnGlyMetSerValGlnLeuHisValProlyrSerGluHisAlaL 	qq	314
		δ	150
	ysAlaProValGlyProValThrMetGlyLeuProLeuProMetAspProLysLySL	qq	317
	CCCCACCGTTGCCGCCAGCAACGGTGAGCCCTCCGAGGGCACCGTGCGCCGCCGTCAC	ò	152
	euAlaProPheSerGlyValLysGlnGlnGlnLeuSerProArgGlyGlnAlaGlyProP	qq	320
	2445 CCTTGCCGCCGTCACCGCCGTCACCGATGTCGCCGGCGTCACCGC 2489	ò	154
0y 1	1181 roGluSerLeuGlyValProThrAlaGlnGluAlaSerValLeuArgGlyThrAlaLeuG 1201		

::: 2490 CCTTGCC	THECETICAGECCAACGGCCC-GCCATCACCGCCGAGAGCTTCG 2536
1201 lySerValProGlyGlySer	IVSerValProGlyGlySerIleThrLysGlyIleProSerThrArgValProSerAspS 1221
	12 26
1241 lyThrIleThrarglleIle :: 2645 GCACCGTTGCC	YThrileThrargileileGlyGluAspSerProSerArgLeuAspArgGlyArgGluA 1261 ::: ::: ::: CACCGTTGCC
1261 spSerLeuProLysGlyHisVallle ::: 2687 CCGTGTTTGCTGGCGTCCACGCCGTT	BpSerLeuProLysGlyHisVallleTyrGluGlyLysLysGlyHisValLeuSerTyrG 1281
1281 luGlyGlyMetSerValThrGlnCysSer ::: :: 2741 CCCCCCAGACCGCCGCGACCGCCTGCGCC	luGlyGlyMetSerValThrGlnCysSerLysGluAspGlyArgSerSerSerGlyProP 1301 :::
1301 roHisGluThrAlaAlaPro	HisGluThralaalaProLysArgThrTyrAspMetMetGluGlyArgValGlyArgA 1321 ggggrCacccccctcccccccccccccccccccccccccccccc
1321 laileSerSerAlaSerile ::: ::: 2828 CCGTTGCCGCCCC	SeralaSerIleGluGlyLeuMetGlyArgAlaIleProProGluArgHisS 1341
1341 erProHisHisLeuLysGluGlnHis 2884 GACCCACCGGTGCCGTGACTGCCGA	SHisLeuLysGluGlnHisHisIleArgGlySerIleThrGlnGlyIleProA 1361
1361 rgSerTyrValGluAlaGl. :: 2914 CCCCGAAGGTGCGGCCCCG	rgSerTyrValGluAlaGluAspTyrLeuArgArgGluAlaLysLeuLeuLysArgG 1381
1381 luGlyThrProProProPr 2974 ACG-CCATCGCCGCCACC	luglyThrProProProProProPerArgaspLeuThrGluAlaTyrLysThrGlnA 1401
1401 laLeuGlyProLeuLysLeuLysProAl	uLysProAlaHisGluGlyLeuValAlaThrValLysGluA 1421
1421 laGlyArgSerlleHisGl	aGlyArgSerlleHisGluIleProArgGluGluLeuArgHisThrProGluLeuProL 1441
1441 euAlaProArgProLeuLysGluGly 	ProLeuLysGluGlySerileThrGlnGlyThrProLeuLysTyrA 1460
1460 spThrGlyAlaSerThrTh :: 3130ACCGGCGCCACC	AlaserThrThrGlySerLysLysHisAspValArgSerLeuIleGlySerP 1480
1480 roGlyArgThr-PheProProValHisPro 	-Phe Proproval His ProLeu AspVal Met Ala AspAla ArgAla Leu 1499
1500 GluArgAlaCysTyrGluG	GluārgālaCysTyrGluGluSerLeuLysSerārgProGlyThrālaSerSerGly 1519
i520 GlySerlleAlaArgGlyAla ::: 3209 TGGCCGAGGGCATCT	rgGlyAlaProValIleValProGluLeuGlyLysProArgGlnSer 1539
1540 ProLeuThrTyrGluAspH	GlyAlaProPheAla

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Qy 2306 IleSerGlnProGlyThrGluIlePhe-AsnMetProAlarleThrGlyThrGlyLeuMe 2325 Db 105226	2442 gThrProLeuThrAsnArgVal-TrpGluAspArgProSerSerAlaGlySerThrProP 2462	Db 105531 CGCCACCACCACCACCACCACCACCACCCCCCCCCCCC

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1644 GCGGGGTGCCGTGCCGCCCCAAGCCGCGGAGGCCTCCCCGCGGGT 1646 AlaAlaAlaTyrTyrLeuProAgHisLeuAlaProAbryPrPPHisLeuTyr 103501 CACCGCGCGTCCCCCCCCCAAGCACACCCGCGCGCTTGCCG 1666 Pro	1925 uaspGlyValTyrProThrLeuMetGluProValLeuLeuProLysGluAl 1942 :::
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104	249	CACACCTUGGTGGCGCCGGCCGCCGCGCGCCACCCTTCCCACC 104293
104	975	OSELLyBG1ySerGluProArgProLeuValProProValSerG1yHisAlaThr11eA1 1995 :::
1	995 348	erPr :: CC
104	2015	oalaseralaserAspProHisargGluLysThrGlnSerLysProPheSerIleGlnGl 2035
104	2035	uLeuGluLeuArgSerLeuGlyTyrHisGlySerSerTyrSerProGluGlyValGluPr 2055
104	2055	uThrHi BAS GCCGCGACC
104	2075	uGluLeuAspLysSerHisLeuGluGlyGluLeuArgProLysGlnProGlyProValLy 2095
104	ഗയ	sLeuGlyGlyGluAlaAlaHisLeuProHisLeuArgProLeuProGluSerGlnProSe 2115
10,	2115	rSerSerProLeuLeuGlnThrAl 2123
10,	2123	aProdlyvalLysGlyHisGlnargValValThrLeuAlaglnHisIleSerGl 2141
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., 01	2161	AlabroLeuTyrSerPheProGlyAlaSer
10	2174	uAspleuArgArgProProSerAspleuTyrLeuPro-ProProAspHisGlyAlaProA 2194
10	2194	laargGlySerProHisSerGluGlyGlyLysArgSerProGluProAsnLysThrSerV 2214
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alrpAs tuberc	9 40 LeuGluTyrGlnHisHisSerArg	Oy 50 AlaSerHisLeuSerProGlySerIle1leGlnProGlnArgArg ::	Qy 65 strain Db 98910	<i>장</i> 점	66	Db 99030 CGGTATCGCCATTCTCGCGGCTGAACGCGAÀAGCGGCTGTAGGTTGTCGATCTCGA Qy 113 LeuProAspProLeuLeuArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySer	78066	Oy 133 GluaspLeuThrLysAspArgSerLeuThrGly	Db 99132 CGTCAAGCCCGGCGAACGCCGCGTAGTGGTCGCGGTGTCGCGCACGCA	Qy 145 LeuGluProValSerProProSerProProHisThrAspProGluLeuGluLeuVal 163 :::	Qy 164 ProProArgieuSeriysGluGluLeulleGlnAsnMetAspArg	Db 99252 AGCGCGTTCGCCGTCGCTCGGTCGGTCACCACCAGAGACTTGTCTGCCAGCAGGCTGA	Qy 179 ValAspArgGlulleThrMetValGluGlnGlnIleSerLysLysLysLysLysGlnGln 198	Qy 199 GlnLeuGluGluGluGluAlaAlaLysProProGluLysProValSerProProProProProProGluLysProValSerProProProProProProGluLysProValSerProProProProProGram		Db 99426 GTCCGGTCAGCAGCGCATGTGACCAGTCGAGGGAAGCCGGCATGGTCT 99473	66	Qy 258 nGlnProSerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLy 278	Db 99531 AGCIGTCGACGATTCGGTCAG	Qy 278 sLysLeuIleLeuTyrPheLysArgArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPh Db 99553	Db 99577 -TGCAAGTTCGATCGCCAGCGGAATGCCGTCGGTGGCAGATCTCGGTCACCA 99632		99633 GGGCGAGGTTGTCGGCAGTGAT	Qy 322ArgargalaLysGluSerLysValArgGluTyrTy 333
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	52786 TGGGGGTCGCCGCGGGGGGACCACCATGGACCGGAGCTGCGCCTGACCG 2151HisbrodindinLeuSerAlabroLeuProAlabroLe	Db 52906 TCACGGGCGAGGGCCGCCTGGAGCGCG-TGTCCGGCACCTGACCCCGGCGGCCACCCGG 52964

2181 52965 CCGTG	SerAspLeuTyrLeuProProProAspHisGlyAlaProAlaArgGlySerPr 2198
2198 0	
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2209 oAsnL 53136 TCGTC	oAsnLysThrSerValLeuGlyGlyGlyGluAspGlyIleGluProValSe 2226
2226	2246
-0	ringgiccgcchcicaccccachcginaccacccacgicciaccciccrist 532
2246 yrArg 53252	yrargaspGlyGluGlnThrGluProSerArgMetGlySerLysSerProGlyAsnThrS 2266
2266 erGlr	ePheSerLysLeuThrGluSerAsnSerAlaMet
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334	
2302 roglu 53385 CCGAC	rogluTyrasnileSerglnProglyThrGlullePheasnMetPro 2317
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2333 lnGl	uHisAlaSerThrAsnMetGlyLeuGluAlallelleArgLysAlaLe
8 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4 4	2369
2353 ysty 53540	 GTC 535
2370P	ProLeuAsnAlaSerAlaSerLeuProAlaAlaMetProlleThrAlaAlaAspGly 2388
53595 TGCC	53
2389 ArgS 53655 CGCT	ArgserAspHisThrLeuThrSerProGlyGlyGlyGlyLysAlaLysValSerGlyArg 2408
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25	2445
53766 CCGT	AGCCGTCCTCCACCTCG
2445 uThrAe 53805 AGGCG	uthrashargvaltrpgluaspargproserseralaglyserthrpro 2461
2462	OTyrAsnProLeulleMetArgLeuGlnAlaGlyValMetAlaSerPr 2479
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Qy 1300 Db 50124 GCAC Qy 1315 uGly Db 50170 Qy 1335 ePhr Db 50178 ECCC Qy 1355 eThr Db 50238 ECCC Qy 1392 pLeu Db 50346 ACCC Qy 1403 Db 50466 CCGG Qy 1423 qSer Db 50466 CCGG Qy 1437 Glu Db 50579 TACT Qy 1437 GLu Db 50636 CACG Qy 1489 OLeu Db 50756 CAGG Qy 1501 Db 50756 CAGG Qy 1501 Db 50816 CCCG Qy 1501<	Oy 1527 Db 50996 GCGT Oy 1530 1 Db 51056 CCAC

seccacicec---- 51785 rThrSerSerSerGl 1795 :::||||||::: cTcGTcTTcGGcGcc 51873 ---CGCGCCCTCC 51900 STCGTGCACTACCTC 51960 rSerArgProAlaSe 1853 | |||||||| | AccGcGcCcCGCA-- 52018 cceecceacecc 51232 ceacecercrece 51292 SerThrProArgGl 1591 |||| ccccrcrccccrc 51538 recerracecees 51595 ccaccecrececk 51655 ----- 51749 |||| cerceaccrecece 51814 ccescccccrrcc 51352 CCAGGCGTA---- 51461 3CGA----- 51494 GGCGCCCTCCCCG 51172 ArgHisSerSerSe 1775 JGluLysSerIleLe 1815 ProArgLeuGlnGl 1571 ProHisProlleSe 1609 SerGlyValAsple 1623 ArgGlylleProLe 1643 Pro-----Th 1660 Thralaalaleu-- 1679 SerGlnGlnMetHi 1696 GlyLeuSerProAr 1716 elleAspLeuSerGl 1736 AlaThrAlaMetAs 1756 ogly---ThrGluGl 1833 ------1555 1------ AB 1869

cccccccccccccccccccccccccccccccccccccc	1218proSerAppSerAlaileThrTyFArgGiySerIleThrHisGiyMirProAlads 1236 19842 CGGACGCCGCCGCCATCGCCTGTCCCCTCGCCCCCCCCCC
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	47115 CGGCCACAGTGCCACGACGACGCGCCCCGAACG	531 LysgluGluGluMysPro	538 ValGluAsnAspLysGluAspLeuLeuLysGluLysThr ::::::::::::::::: 47234 GCGGATGATGAACGACCGCACCGCACCGTGCCGGCGCACA	551AsphapThrSerGI 47294 GACGCCGCGCGCACGCACGCACGCCGGGCTCTGGCACTGGTGG	562 GluAlaValAlaSerLysGlyArgLysTh ::: 47354 CCACACAAACACACACACACACACACACACACACACAC	578	47414 CATCCCGGGGACCGGGTCGCCCTCGCCGTCGAGTACGGCT	579 LysclyargliefnrargsermetAlaAsnGluAlaAsnse 47474 CCTGGCGCTCCGCGCGGAGCCGTCTGCCTGCCGTCG	598 oGlnGlnSerAlaG 	4.734 Gettigeraseraterioresectessectes	::: 3.1 47594 CACCCAGGATCGACTGGCCGCAGGAA	629 uHisGlyArgAsnTrpSerAlaIleAlaArgMetV	4/836	47695 CGACCGGCCTACCGGGTGGACGCCCCCGTCAGCCAC	667 nGlnHisbysLeubysMetGlubysGluArgAsnAlaArgA	687 OAlaAlaSerGluGluAlaAlaPheProProValValG		707 aSerGlyValSerG	TTCGGGCCCCTCCTGGCCGGTGC	/IB IGIUGIUAlaguwiaku :	732 OArqGlvGluCysSarGlyProAlaThr		742ValhanAsnSerSerhapTi	4002/ CICCGGCIGGIGGIGGCIGGCGGCGGCGGAACGCCICGGCGGCGGCGGCGGGGGGGG	
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.52386
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46218 CCGCGC------CACGCCGAGG 46235 -----ccceecceecrieccerecesteceecce 46157 |||::: :::||| 46008 GGTGCGCGACGCTGCTGGTTCGACGTACGGACCCA-----CCGCCTGGA 46061 117 LeuLeuArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySerGluAspLeuThr 136 97 LysSerGluMetGluPheIleGluSerLysArgProArgLeuGluLeuLeuProAspPro 116 26 96 The present sequence represents the BLM (Bleomycin) gene cluster, containing open reading frames (ORFS) 8-30. The proteins encoded by the gene cluster are useful for producing peptides and/or polyketide metabolites, especially bleomycin or bleomycin analogues. They are also useful for chemically modifying biological molecules to produce branched methyl groups, and for coupling amino acids and fatty acids. They are reacted with an apo-carrier protein and coenzyme A to produce a holocarrier protein. The BLM gene cluster or catalytic domains can be used individually or collectively to produce thiazolidine, thiazoline, bithiazoline and bithiazoline-containing microbial metabolites. The BLM gene cluster may also be used to produce sugars 3 GlySerThrGlnLeuValAlaGlnThrTrpArgAlaThrGluProArgTyrProProHis 22 41 GluTyrGlnHisHisSerArgAspTyrAlaSerHisLeuSerProGlySerIleIleGln 77 GluargSerGlnGluLeuHisLeuArgProGluSerHisSerTyrLeuProGluLeuGly 137 LysAspArgSerLeuThrGlyLysLeuGluProValSerProProSer------23 SerLeuSerTyrProVal-----GlnIleAlaArgThrHisThrAspValGlyLeuLeu P-PSDB; AABO7556, AABO7557, AABO7558, AABO7559, AABO7560, AABO7561, AABO7562, AABO7563, AABO7564, AABO7565, AABO7569, AABO7569, AABO7579, AABO7571, AABO7572, AABO7573, AABO7574, AABO7575, AABO7576, AABO7577, AABO7578, Sequence 58857 BP; 7256 A; 25139 C; 19353 G; 7109 T; 0 U; 0 Other; New bleomycin gene cluster components useful for peptide and/or polyketide metabolites, especially bleomycin, production and for chemically modifying biological molecules. |||| 46062 CGACGTCGGACCCGGCTGCACCTGAGCTTGGACCTGGTCAT----58857 681 275 1098 1078 Length: Matches: Conservative: Mismatches: Edwards DJ; Indels: US-09-522-753-5 (1-2517) x AAA58471 (1-58857) Claim 8; Page 97-136; 162pp; English. Chen M, 2.29e-11 666.00 30.64% 21.83% 5.04% Sanchez C, CIGGCGGGCCCGGCI UNIV CALIFORNIA WPI; 2000-465974/40. Percent Similarity: Best Local Similarity: Da 15, Alignment Scores: 46104 Best Local Si Query Match: DB: (REGC) Shen B, No.: Score: Pred. g à 쉽 ò ò 9 ð g ò 음 ð g à 셤

ò	1311 TyrAspMetMetGluGlyArgValGlyArgAlalleSerSerAlaSerIleGluGlyLeu 1330	
q	98250 GCCGGCGCCGCCGACGGTGATACCAATGCCGCCGTTGCCGCCGGCCCCCGCCAACGCC 98306	43014
λō	1331 MetGlyArgAlaIleProProGluArgHisSerProHisHisLeuLysGluGlnHisHis 1350	QY 1082 rgGininiieilei
qa	98307 GCCGGCCGCCGCCGCCGCCGCCGCCACCGGTGCCGTGACTGCCGA 98362	
ò	1351 IleArgGlySerIleThrGlnGlyIleProArgSerTyrValGluAlaGlnGluAspTyr 1370	אַר פּסָטעַט אַר אַר
셤	98363CCGTCCCCGAAGGTGCGGCCCCGACCCCAC 98395	
ογ	1371 LeuArgArgGluAlaLysLeuLeuLysArgGluGlyThrProProProProProPeroSer 1390	
q	98396 CGTCCCCGCCATGTCCACCGACCCCGCCGCACCG-CCATCGCCGCCACCACCACCGCCGCC 98454	TAC 1745
ò	1391 ArgAspLeuThrGluAlaTyrLysThrGlnAlaLeuGlyProLeuLysLeuLysProAla 1410	OY 1/42 Albenval-Fiores
qa	98455 CCACCGGIGCCGCCGAIACIGICGAIACCGIIGGCGCCCCCIG98496	(711)
δ	1411 HisGluGlyLeuValAlaThrValLysGluAlaGlyArgSerIleHisGluIleProArg 1430	DY COUNTRICTORY
đ	98497GCCCCGGCC 98505	
λō	1431 GluGluLeuArgHisThrProGluLeuProLeuAlaProArgProLeuLysGluGly 1449	99208
q ₀	98506 ccaccectáececcacacacertáceceseces	1802 ArgGluA
ò	ω·	99237
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ò	1489 oLeuAspValMetAlaAspAlaArgAlaLeuGluArgAlaCysTyrGluGluSerLeuLy 1509	2501
අු	98631 C 98631	7 2 2 2 1
ò	1509 sSerArgProGlyThrAlaSerSerSerGlyGlySerIleAlaArgGlyAlaProVall1 1529	O.
ąg G	98632 -GCGTCGCCCTGACCCCCGTTGCCTCCCGAGGCCGAGGCGATCT 98682	7,000
ò	1529 eValProGluLeuGlyLysProArgGlnSerProLeuThrTyrGluAspHisGlyAlaPr 1549	99431
ΩP	98683GAACCAGGCGCGCCCGAATTGGCCCCGCCGCCGCCGCCGCCATCGCCACC 98735	1886
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qq	98736 ATrecedececeaceacaceaccaceaccedeceacarraceacere 98783	1906
ò	$r ProArg Leu Gln Glu Gly Ser Leu Ser Ser Ser Lys Ala Ser Gln Asp Arg Lys Leu Th. \\ \ \ \ \ $	99523
අ	GCCTCCGTTGCCCCC	Qy 1926 AspGlyValTyrPr
ð i	rSerThrProArgGlulleAlaLysSerProHisSerThrValProGluHisHisProHi	Db 99583 CCGCCGTTACCACC
g G	GEOGCOGCOCGOGACGCTGGCTTTGCGCACCGTTTACCACCGTTTGCCGCCGCT	Qy 1943 ProArgValAlaAr
ð í	BProlleSerProTyrGluHisLeuLeuA :::	Db 99643 CCGTTACCGCCTTV
<u>a</u> d	GCC6GGCCCG1GG	Qy 1956 GlyHisAlaPheLe
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3	ACCOCCE 1.000000000000000000000000000000000000	Qy 1996 ArgThrProAlaLy

hrLysProThrThrSerSerSluArgGluArgAspArgAsp 1801 ||| ||||||||||||| |CGCCGCCGCC-ACCACTAGCTCCGGC------99236 cadecacedecerracecerredececenedareceredece 99642 SCCGCGCCGC-CGGCCTGCCTCGACCGCGTTGCCGC 99370 || :::||| :::||| Traacceccarrecccearreccccaaage 99490 ...||||||||||... recciccerreceaceacrerecacegraececcicegrecicecere 99702 cecceccicicicede CACCATCGCCGCCCCCCCCG-----TCACCATCCTTGAAAGCC 99801 ----- 99116 galaaspMetLeuargGlyLeuSerProArgGluSerSerLeuAlaL 1722 leTrpArgProGlyThrGluGlnSerSerGlySerSerGlySerSer 1841 euGlnGlnArg---ProSerValleuHisAsnThrGlyMetLysGly 1885 alGluProSerLysProThrValLeuArgSerThrSerThrSerSer 1905 laalaThrPheProProAlaThrHisCysProLeuGlyGlyThrLeu 1925 roThrLeuMetGluProValLeuLeuPro-----LysGluAla 1942 rgProGlu------1955 euAlaLysProProAlaArgSerGlyLeuGluProAlaSerSerPro 1975 luProArgProLeuValProProValSerGlyHisAlaThrIleAla 1995 eAsnAspTyrIleThrSerGlnGlnMetHisHisAsnThrAlaThrA 1702 aGlyProArgGlyIleIleAspLeuSerGlnValProHisLeuProV 1742 roThrProGlyThrProAlaThrAlaMetAspArgLeuAlaTyrLeu 1761 InProPheSerSerArgHisSerSerSerProLeuSerProGlyGly 1781 rgAspArgGluArgGluLysSerIleLeuThrSerThrThrTval 1821 ----SerHisAlaHisGlnHisSerProIleSerProArg----- 1866 ysasnLeualaProHisHisAlaSerProAspProProAlaProPro 2015 -ProTyrLeulleArgGlyTyrProAspThrAlaAlaLeuGluAsnA 1682 |||||||| CAACGCCGCCGACACCGCCGT----cccecc-gracecccaacgccgccaacaccacacacca lySerSerArgProAlaSerHis-----

298 eCysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGluArg1leGluAs 318	338 oGlulleArgLysGlnArgGluLeuGlnGluArgMetGlnSerArgVa 354	7 354 1GlyGlnArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluHisGluValSe 373	373 rGluIleIleAspGlyLeuSerGluGluGluAsnLeuGluLysGlnMetArgGlnLeuAl 393	393 aVallleProProMetLeuTyrAspAlaAspGlnGlnArgIleLysPheile 410	/ 411 -AsnMetAsnGlyLeuMetAlaAsp	/ 421 sValTyrLysAspArgGlnValMetAsnMetTrpSerGluGlnGluLysGluThrPheAr 441 :::	/ 441 gGluLysPheMetGlnHisProLysAsnPheGlyLeuIleAlaSerPheLeuGluArgLy 461	461	/ 481 rLeuValargargSerTyrargargargdlyLysSerGlnGlnGlnGlnGlnGlnGlnGl 501	501 nGlnGlnGlnGlnGlnGlnGlnGlnBroMetProArgSerSerGlnGluGluLysAs 521	/ 521 pGluLysGluLysGluLysGluAlaGluLysGluGluGluLysProGluValGluAsnAs 541	/ 541 pLysGluAspLeuLeuLysGluLysThrAspAspThrSerGlyGluAspAsnAspGluLy 561	/ 561 sGluAlaValAlaSerLysGlyArgLysThrAlaAsnSerGlnGlyArgLysGlyAr 581 ::: 96223CGACGCCGGGGTGA 96236	581 gileThrArgSerMetAlaAsnGluAlaAsnSerGluGluAlaileThrProGlnGlnSe 601	601 rAlaGluLeuAlaSerMetGluLeuAsnGluSerSerArgTrpThrGluGluGluMet 620	621GluThrAlaLysLysGlyLeuLeuGluHisGlyArgAsnTrpSe	635 rAlaileAlaArgMetValGlySerLysThrValSerGlnCysLysAsnPheTyrPh 654
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Pred. No.: 3.1e-11 Length: 110000 Score: 668.50 Matches: 617 Percent Similarity: 32.38\$ Conservative: 255 Best Local Similarity: 22.91\$ Mismatches: 1044 Query Match: 5.06\$ Indels: 793 DB: 4 Gaps: 113	4 SerThrGlnLeuValAlaGlnThrT	Qy 24	37 ValGlyLeuLeuGluTyrGlnHisHisSerArgAspTyrAlaSerHisLeuSerProGly	57 SerilelledinProdinArgArg-ArgProSerLeuLeuSerGluPheGlnProdiyAs [76 nGluArgSerGlnGluLeuHisLeuArgProGluSerHisSerTyrLeuProGluLeuGl 	96 yLysSerGluMetGluPheIleGluSerLysArgProArgLeuGluLeuLeuProAspPr	95147	95170 CCAGCGGGGAACCCGCCAAATCTGCTCACCGGCCAC	156 rAmpProGluLeuGluLeuValProProArgLeuSerLysgluGluLeuIleGlnAsnMe	Lysleulyslysly :::::::::::::::::::::::::::::::	196	216 oProProlleGluSerLysHisArgSerLeuValGlnIleIleTyrAspGluAsnArgLy 95341 TAAGAGAAATGAAATAAAAAAAAAAAAAAAAAAAAAAA	236 sLysAlaGluAlaAlaHisArglleLeuGluClyLeuGlyProGlnValGluLeuPro	95401 GCGCCGCGCGATCGGCACCACCAGCGCCGGGTCGGTCGGCGCCCAGGTTCGACATACCAAA 256	OY 258 nGlnProSerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLy 278 Oy 258 nGlnProSerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLy 278 nh eses: carcractarfactarasparasparasparasparasparasparasparas	278 slysleulleLeuTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPh	

1899	49747 CGCCGCCGCCTGCATHACTAGATAACCCTCGAAACCCTCGGCCCCTGGTGACGA 49688 2121 lnThrAlaProGlyValLySGlyHisGlnArgValValThrLeuAlaGlnHislle 2139 [
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1684	CCGGGAAGAAGCCAACGAAAGCGCGCATCAAGGGATGAAATATCGGGACGATGATATCCAA

ov 827 vsGluGluGluTbrAlaAla	::	834	74 A A	8 73	54466	873	54411 TGCCGGCAGCCGGGCTCCCAGCAC	Qy 889 lyAlaLeuLysAlaGluLysLysGluGlyGlySer	54	Qy 909 erGlyAlaProGlnAspSerAspSerSerAlaThr	Db 54307 GGGCGCACAGATACCCCAGCACCTCGCGGCG	Qy 929 laGluGlyGlyAspLysAsnArg	Db 54247 dfrctrcgrccccaaggrcagraccacracracac	Qy 941 ArgProSerLeuLeuThrProThrGl	Db 54187 GTGCCGTCAGCATCGCGGGTAGCGTTCCCGACCGG		54127	QY 9/3 IIeProtroinedinvalini		UY 54. FIOF UNIGGIANS PARAMETER IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT IN INTERPORT INTERPORT INTERPORT INTERPORT IN INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPORT INTERPO		'n		53887 AATTACCCTGACCCGCGCGCCC	Qy 1025	Db 53827 ACAAGTCCATATCCCGAGTCAGCTCGTGCAGGTTC	Oy 1033 GluAlaPheAlaAlaGluAlaGlnLysLeuProGl	Db 53767 ACACCGCCCCCC	Qy 1053 LeuproPheProValProProArgGluVall1	Db 53731 TCACCCGTCAGCCAGCACCCGCGG	Qy 1072 ProSerAlaPheSerTyrAlaProFroGlyHisPr	53692	1090	Db 53638 ĠATCCGCAACGTCGCACGCCACCACCATGACCACC Qy 1107 ProLeulleSerSerAlaLySHisProSerValLe
	515SerSerGinGluGluLyBApgluLyBGluLyBGluLyBGluLyBGluLyBGluLyBGluGluGluG	55398 GATCACCCTTTGCTGCGAGGGACCATTCGGCGGTCAGTCCGTTTGACGCGCCGTCCTG 55339	534 luLysProGluValGluAsnAspLysGluAspLeuLeuLysGluLysThrAspAspThrS 554	55338 ATTCACCGCCGACCCACGAACCACCGCCAACACCCCGATG 55300	554 erGlyGluAspAsnAspGluLysGluAlaValAlaSerLysGlyArgLysThrAlaAsnS 574	55299 CCCATTACGACGGCATCCGACAACCTCTCCAACAGCACTAGGCCAGCACCCTCGCCCCA 55240			luLeuAlaSerMetGluLeuAsnGluSerS	CTGACGGGAGAACTCCACAAACGTCGCCG			633 BNITDSETALIALIEALANGMECVALGLYSEILYBINIVALSEIGINLYBHYBHBURNEI 955 :: :	nAsnteuAspGluIlebeuGlnGlnHisbysbeubysM	55129	etGluLysGluArgAsnAlaArgArgLysLysLysLysAlaProAlaAlaAlaSerGluG		693 luAlaAlaPheProProValValGluAspGluGluMetGluAlaSerGlyValSerGlyA 713		713 snGluGluMetValGluGluAlaGluAlaLeuHisAlaSerGlyAsnGluValProA 733	54994CCGA 54991	733 zgGlyGluCysSerGlyP 739	54990 GGCCACACTGCCGCGCTGCCATTACCGAGATAGCCTTCGAAGCCCTCCGGCACTCCGGC 54931	739 roAlaThrValAsnAsnSerSerAspThrGluSer1leProSerProHisT 756	ပ္ပ		GCGCAGCGACCCCGGATCCCCAGCGCGTTCAACGGCCTCCCAGGACAGCTCCAGCAG		CAACCGTTGCTGCGGGTCCATCGCCAGTGCCTCACGCGGAGAAATCCCGAAAAAGGCGGC	783ProThrProProArgArgThrSerArgAlaPro		794IleGiukTountkrohlaserGiualaintotyalarroi ao/ 		
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cgrdcrcccaccacccaagcaagg 54248 ACGCCAGCCCGGCATCGAACAGCCGCA 54068 GGGCGATCCTGCTCCGGTCGACATCAC 54008 :::|||||||::: cgrccaggaacgrgrrcgccgccar 53888 ACAGCGAGGAGAACAACGCGAAAAACG 53828 | | | | cGCCAGCAACCCTCCAGACCAGCAC 53639 CCGCACCGAAACCCTCCAGCTCACCCA 53579 SGCCGTCGGGTAGTCGAATACGGCGGT 54412 |||||::: GCCTGATAGAGGTGATGGTAGGTACCA 54188 3GGCCTGCTCGGCCAGGAAGGCCCTGT 54128 ACAAGCCCCACGCCAACGACACCGCAG 53948 aGlualaalaGlualaThralaGluG 889 ||::: ||| :::||| :::||| ::GATTACGCAAC-TCCACACCAGACA 54353 rGlyArgAlaThrThrAlaLysSerS 909 SAGATCGTCTTCGACGCTCGCGGTCGT 54511 3CCTGCCAGTT----- 54467 ---SerProAlaProProAlaAspLys 1032 GlyAspproProCysTrpThrSerGly 1052 IleLysAlaSerProHisAlaProAsp 1071 Pro-----LeuProLeuGlyLeuHig 1089 ThrileSerAshPro-----Pro 1106 LeuGluArgGlnIleGlyAlaIleSer 1126 AlaProProAlaProProProGln 1003 SlnProglySer----- 1017 3lyLysSerArg----- 1024 erGluCysThrGluGluAlaGluGluG 873 hrCysserAlaAspGluValAspGluA 929 rgreuren-----Ser-Pro 940 JyAspProArgAlaAsn----- 954 LysGlnLeuLysGlnArgAlaAlaAla 972 ------bysValHisGlu 983 ysproproAlaAlaGluGluLeuAlaV 853 .----- 833

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that encodes a butenyl-spinosyn biosynthetic enzyme, a butenyl-spinosyn RS (polyketide syntheties) domain, or a spinosyn PKS andule. The butenyl-spinosyn biosynthetic genes are useful for increasing the production of butenyl-spinosyn insecticidal macrolides. The genes are also useful for fanging the metabolites or products produced by spinosyn-producing microorganisms. The present sequence represents a DNA molecule encoding butenyl-spinosyn biosynthetic enzymes
                                                                          the
changing
                                                                                                                                                                                                                                    invention relates to a novel DNA molecule comprising a DNA sequence
P-PSDB; ABP57678, ABP57679, ABP57680, ABP57681, ABP57682, ABP57705,
ABP57706, ABP57707.
                                                                        New butenyl-spinosyn biosynthetic genes, useful for increasing production of butenyl-spinosyn insecticidal macrolides, or for the metabolites or products produced by spinosyn-producing microorganisms.
                                                                                                                                                                                        Claim 2; Page 57-99; 218pp; English.
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ž S	Sequence 75236	75236 BP;	10935 A; 21693	C; 28185 G; 14421	21 T; 0 U; 2 Other;	
Alignm Pred. Score: Percen Best L Query	ent No.: t Si t Si Matc	Scores: milarity: Similarity: h:	2.23e-11 669.00 30.72\$ 21.47\$ 5.06\$	Length: Matches: Conservative: Mismatches: Indels: Gaps:	75236 694 299 11134 153	
us-o	9-522-753	-5 (1-2517)	US-09-522-753-5 (1-2517) x ABV75557 (1-	(1-75236)		
ઠે	ı	MetSerGly8	SerThrGlnLeuValA	MaginThrTrpArgA	MetSerGlySerThrGlnLeuValAlaGlnThrTrpArgAlaThrGluBroArgTyrPro 2	20
셤	56835		GITTCCGGTCAGCGTCGAGACAATCGGCAGCCCTGGAGACCGGT-AATCCACGC	GGCAGCCCTGGAGAC	-	56783
à	21		seuSerTyrProVal	inileAlaArgThrH	ProHisSerLeuSerTyrProValGlnIleAlaArgThrHisThrAspValGlyLeuLeu	40
g	56782		TCGTGC	GATGTCGGCGAACTC	TCCGTGCGATGTCGGCGAACTCATCC-AGCATCGGATCCATC	56742
ઠે	41	GluTyrGln	HisHisSerArgAspT	YrAlaSerHisLeuS	GlujyrGlnHisHisSerArgAspTyrAlaSerHisLeuSerProGlySerIleIleGln (09
g	56741	CGATACGAAT	6741 CGATACGAATGGAACGCATGCGAACCGCAACCAT	. : ! .ccgcaaccat		56706
Š	61		ArgArgProSerLeui	euSerGluPheGlnP		80
g	56705		GGAACACCCAGC		CGCCACCGAACACCCAGC	56679
à	81		ceuArgProGluSerF	lisserTyrLeuPr	SerGluM	100
g	56678		CGGCCAACAACGTCCTCAAGCACATCCCGGTGACCGGAAAG	CCTCAAGCACATCCC		56638
ò	100		leGluSerLysArgPr	oArg		114
ą G	56637		CACCGCCGACC	 CCGGAGCATTAACCG	CACCGCCGGAGCATTAACCGCTGCGACACAACACCGATCC :::	56592
ò	114		euLeuArgProSe	rProLeuLeuAlaTh	roAspProLeuLeuArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySerG	133
ą a	56591	CGCACCCCAT	:: rccagcagcgggctgA	rrcggreeregereg		56533
à	133		hrLysAspArgSerLe	uThrGlyLysLeuGl	luAspLeuThrLysAspArgSerLeuThrGlyLysLeuGluProValSerProProSerP	153
e G	56532		GGCAACGCCTGCATC	AACCGGGCACGACCG		56473
à	153		hrAspProGl	.uLeuGluLeuValPr		171
g	56472		I I CAGCACACCGGCGGCG	SAACGCCGCTGCTAGC		56413
à	171		InAsnMetAspArgVa	lAspArgGluIleTh	luLeulleGlnAsnMetAspArgValAspArgGlulleThrMetValGluGlnGlnIleS	191
g G	56412		AACAGGCCGAACACC	CACGAACCCAACAAC		56356
ઠે	191	erLysLeuL)	erbysLeulysLysLysGlnGlnGlnLeuGluGluGluAlaAla	InLeuGluGluGluAl	LysP	207
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CGTCGGAACCGA	sHisArgSerLe cgccacaAgcT7	sArglleLeuGlu	pThrargglnT) cacccTGaccGC	euTyrPheLysArg ::: CACAGATCC	nTrpLysGlnLy ::: :: CGGACAACAACC	uLysLysValG] :: GCGTCATCGCC	ggluTyr ATCGATCGTCC	-TyrGluLysGlnPh ::: mrrrcccacacaca	gValGlyGlnAı 	r : TTGACGCTGTCC	nLeuGluLysG] :: ACGACACTCCC	aAspGlnGlnAı ::: GCTGGACCGAAG	alTyrLysAspArg	uLysPheMetG] ::: TCACCATCTTG	rvalAlaGluC, ::: CCTGTC	euvalarg-arg GATCACGATCA(GlnGlnGlnGlr ::: TGGTCCCGGTCC	TGAGTGCTTGGC
CGCGAACAAACCCGACTGCGCCCACAACGTCCGATCGAGCAACTGCGCGTCGGAACCGAA	roproglubrogluLysProValSerProProlleGlu-SerLysHisArgSerLeu 	aGluAlaAlaHi	GlyLeuGlyProGlnValGluLeuProLeuTyrAsnGlnProSerAspThrArgGlnTyr	- er	ArgasnHisalaarg	PhecysglnargTyraspGlnLeuMetGlualaLeuGlubysLysValGlu	ArgileGluasnasnProargargalaLysGluSerLysValargGluTyr 		eArgLysGlnArgGluLeuGlnGluArgMetGlnSerArgValGlyGlnArg 	erMetSerAlaAlaArgSerGluHisGluValSer		MetargGlnLeuAlaValIleProProMetLeuTyrAspAlaAspGlnGlnArg :::: ::: ::: ::: ::: ::: ccGAcGATGGCCGTGTCGGGCGTGTTCTCCGTGAGGAGGACGGAC	pPrometLysVa CGTG	alMetasnMetTrpSerGluGluGluLysGluThrPheArgGluLysPheMetGln::: ::: :::: caarGTGCGCGCAGCTCCCCGTGGCGCATCGCCATCTTGAT	HisProLysAsnPheGlyLeuIleAlaSerPheLeuGluArgLysThrValAlaGluCys 	ValleuTyrTyrTyrTyrLeuThrLys-LysAsnGluAsnTyrLysSerLeuValArg-ArgS 	ertyrargargalylysserginginginginginginginginginginginginging	InglnglnglnglnprometProarg
CAACGTCCGAT	SerProProPro	Valginileiletyraspgluasnarglyslysal. ::: ::: GCCAAAGGCATCAGCGAACACCGGAAAGGC	1ProbeuTyrAs: CCGGCCCATCC	eLysileAsnGlnAlaMetArgLysLysLeuil 	BACCGGCAGCCA	CASPGInLeuMe	ArgAlaLysGl		ileuGlnGluar :::: DATTCACAGCAG	AAlaArgSerGl GTGCTC	AspGlyLeuSe ::	oProMe sccAgggcgrgr	IleLysPheIleAsnMetAsnGlyLeuMetAlaAspProMetLysV TGCCGACCATCCACTCGACGTGGGCTCGCCGACGTG	JGlnGluLysGl JCAGCTCCCCGT	eAlaSerPheLe	s-LysAsnGluA ccccAaccaca	slnglnglngln ::: stgcctcGATCG	arg GGACACGGACA
CGACTGCGCCCA	luLysProvals AccccactTC	fyraspgluasr ::: agcgaacac	31nValGluLer ::: ccGTAAAGCCC	Lys I leAsnGlr AAACCGGGCCC	Arg	GlnArgTyr ::: cccAGCACCAC	AsnProArgArg ATCCCGCGA		LysGlnArgGlu ::: AACCACAGGGAC	SerMetSerAla TCCGCT	GluIleIle ::::: SATGACGTGTGC	AlavalilePro ccGcTGTCGC	AsnMetAsnGly :::::: CACCTCCGACG1	MetTrpSerGlu ::: CAATGTGCGCGC	PheGlyLeuIle	TyrLeuThrLye GGACTTCATCG	rgGlyLysSerC GATCAATGCCTC	InProMetPro/ GCATCCACAT
GCGAACAAACC	oProGluProG. ACCACATCCCG	alGlnIleIle: ::: TCAAAGGCATC	lyLeuGlyPro(HisGluAsnIlel GAACAC	rgAsnHisAla) GTAACCACCC	heCys TCACGGTCCGA(rgileGluasni GaataCCCaaCi		oroglullearg	lySerGlyLeu	GACTGTTCGAGG	letArgGlnLew :::::: :cGACGAGGATG	leLysPhelle ::: GCCGACCAATC	JnV	tisProLysAsni 	alLeuTyrTyr' iTGACCGATGTT	rTyrArgArgA: ::: ccatAGGTGGC	InGlnGlnGlnProMetProArg- :: grgcgccrccaCaGarccaCarcGGa
56355 C	207 r 56295 C	227 V 56235 G	247 G	267 H 56157 G	287 A 56118 C	298 F 56058 T	315 A 	333 -	338	358 G 55851 -	374 - 55815 A	389 M	407 I 55695 T	427 G 55656 -	447 F	467 V 55578 G	486 e 55518 G	506 1 55458 G

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1226	gGlySerIleThrHisGlyThrProAlaAspValLeuTyrLysGlyThrIleThrArg11 1246
1246	elleglygluaspSerProSerArgLeuAspArgGlyArgGluAspSerLeuProLysGl 1266 :::
266	HisValI ::: AAC
286	31nCysserLysGluAspGlyArgSerSerSerGlyP CCACCAAGAGAGCGAGGTCCCCACGCC
1306	ProLysArg
1326	uG]
1338	GluàrghisSerProhishisLeuLysGluGlnhishisIleArgGlySe 1354
1354	rileThrGlnGlyIleProArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGl 1374 :::
1374	ualalysleuleulysargGluGl 1382
1382	yThrPro
1392	pleuThrGluAlaTyrLysThrGluAlaLeuGlyProLeuLysLeuLysProAlaHisGl 1412
1412	uGlyLeuvalalaThrValLysGluAlaGlyArgSerIleHisGluIleProArgGluGl 1432
1432	uAlaProArgProLeuLysGlu cGGCCCCGC
1452	rGlnGlyThrProLeuLysTyrAs
1472	pvalArgSerLeulleGlySerP : ::: GCCACAAGGGGTC
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1527	oValileValProGluLeuGlyLysProArgGlnSerProLeuThrTyrGluAspHis 1546

7268 CCGCGAGGAGCGCCGCCAGTGCGTCCCGGTCG		669HisbysLeuLysMetGlubysGlub	686 AlaProAlaAlaAlaSerGluGluAlaAlaPh	1098 GCGACCTGCGACCCAGGCACCGGT	704 GluMetGluAlaSerGlyValSerGlyAsnGl	7044 GACCACCGGTCGTCCGGCACGACCACGTCCGA	724 LeuHisAlaSerGlyAsnGluValProArgGl		743 AsnAsnSerSerAspThrGluserIleProSe 			778 ProproProGlyProProThrProPro	6809 CCGCCAAGCCGCCCCCCCCAGTTCG	795 GluProThrProAlaSerGluAlaThrGl	6749 GCTGAGCGCCCCGACGGAAGGGGTCTCGTCC	811	6689 TGATCCAGAGCGGGGCTTCGGGTACCGCCTCG	817 AlaProProProValValProLysGluGluLy.	6629 CGGCGCGCCTCGGCCTGGGCCAGGCCAAC	6570 TCACCGGCGCGCAACAACATCCCCGCGGCTGCT	857 LysAlaGluGluProValLysSerGluCysTh	6516 ACGCCGACGCTGTGGACCTCGGCCCC	877 GlyLysAspAlaGluAlaAlaGluAlaThrAl		897 GluglyGlySerGlyArgAlaThrThrAl	6405 GCAGGCCCGGGTTGCCGCGGTCGACGACGGC	907	6345 CCGGCCGTCGACCGTCCCCGCTCAGTGGC	915 Ser	6285 TCGACGCCGGTCACGGCGGTGCGAACGGGGTC	6225 TAGGTGGGAGGTCGATGCGCCGGCCGGTGCT	
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937 CATACACCACCACCACCACCACCACCACCACCACCACCAC	à à	rPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPheCysGlnArgTyrAs	ପୁପ	 7517 CGTCGAGCGCGGCGTTGGCGGCCGCGTA
### ### ##############################	8 8	CAIAGETCTIGECCGGGCACCCCCGG	ò	
323 gAlaLysGluSerLysValArgGluTyrTyTGluLysGlnPheProGluLlaArgLysGl 343 844	- 원	carccaccaccaccaccaccacacacacaccaccaccacc	셤	
### ### ##############################	ò	gAlaLysGluSerLysValArgGluTyrTyrGluLysGlnPheProGluIleArgLysGl	ò	
343 nArgGluLeuGlnGluArgMetGlnSerArg	d d		අු ,	
### ### ##############################	ò	nArgGluLeuGlnGluArgMetGlnSerArg	ð í	
354 -ValGly	qq		g &	
### ### ##############################	ò	-ValGlyGlnArgGlySerGly	3 8	
366	QQ	cárcadecaranceacarádadaseas acestas castes es contra estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas estas es	ò	
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380 rGludGludAsnLeadGlubysGlubear	д	ACAGTTCGTCGTGCAGGTGGGAGGCGAGAGGGGGGGTCGGGTCGAA	ð	
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393	qq		ò	669HisLysLeuLysMetGluI
### ### ### ### ### ### ### ### ### ##	ò	etLeuTyrAspAl	qq	992292
402 aAspGlnGlnArgileLysPheileAsnMetAsnGlyLeuMetAlaAspProMe 420	g	45G	č	
8060 AGCCCAGCTCCGGCTCGACCGCGCGCGCGCCCT 8004 420 LIysValTyrLysAspArgGlnValMetAsnMetTrpSerGluGlnGluLysGluThrPh 440 8003 CCGAGCGCACAACGCCACACGCCCACGCGCACGCGCTT 7968 440 eArgGluLys	ò	eLysPheIleAsnMetAsnGlyLeuMetAlaAspProMe:	q d	
420 tlysValTyrLygaspArgGlnValMetAsnMetTrpSerGluGInGluLysGluThrPh 440 8003 CCGAGCGACAAGGACACACACACACACACACACACACAC	q	CGACCGCGTCGGCCGAGGCGTGGCGCAGGATCCCGGCGGCCT	ò	
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:::|||| ;rccaaggrgrcgrrccgrgcgrccrcgccgacg 6985 decedentacesas decesas da de 1758 TCTCGCCGGCGTGCGTCCCTGCGGCAGAGG 7698 AGGACGTCGAGCGCGCTCGGGGTCCAGGGCCG 7638 AGACGGCCTGCCTCCGTTGCCCCACACGCCGA 7458 CGGTCGGAGGTGAGCGAGTCGAGC-----ACAC 7329 GGTCGCCACGTCGCAGGCCTCGACGGC-CACT 7210 ACCACTCCCCACTCCGGGCCGGCCGCCCG 7150 |||| ||||| :::||| |CGAGCGGAGGCGGGACGGGGACGGGGGCGGGG- 6926 GGAGGCCGTACCGCGAACTGGTTCTCTCCCCC 6869 GGAGCACCGCGCCACTCCCGGCCAGGT 6810 scoccorricidadeccarrecocacedece 7578 || ||||||| |CGAGCGGAAGGTCCGCGGTCAGCTCCAGGT 7398 -----GCGGCGTGA 7383 rgargSerTyrArgargArgAlyLysSerGlnG 495 ingln-----Glnglng 509 luGluLysAspGluLysGluLysGluA 529 ysGluAspleuLysGluLysThrAspAspT 553 ------GluA 557 serLysglyArgLysThrAlaAsnSerGlnGlyA 577 MetAlaAsnGluAlaAsnSerGluGluAlaIleT 597 serMetGluLeuAsnGluSerSerArgTrpThrG 617 31yLeuLeuGluHisGlyArgAsnTrpSerAlaI 637 --LysThrValSerGlnCysLysAsnPheTyrP 654 AspGlulleLeuGlnGln------688 AsnGluGluGluMetValGluGluAlaGluAla 723 ArgGlyGluCys---SerGlyProAlaThrVal 742 ProSerProHisThrGluAlaAlaLysAspThr 762 -----ProProAlaThrLeuGlyAlaAspGly 777

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                                                                                                                                                                                                                                                        20277 GCAACGACTCCTCGAAACCCTGGGAAACCATCGAACACGCCGGCATCAACCCCCA 20336
  20097 GGACGCGATCGGCGGATTCCCCACCGACCGCGCTGGGACCTGGACACGCTCTACGACCC 20156
                                                                                                                                   2483
                                                                                                                                                                                                                            2483 lyLeuProAlaGlySerGlyProLeuAlaGlyProHisHisAlaTrpAspGluGluProL 2503
                                                                                                                                                                           20217 CCACTTCGACGCCGAATTCTTCGGCATCAGCCCCCGCGAAGCCCCTCGCATGGACCCCCA
                                                                                                                                   -----IleMetArgLeuGlnAlaGlyValMetAlaSerProProPro-----ProG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Tylactone synthase gene cluster; tylG gene; multifunctional protein; platenolide synthase gene cluster; platenolide production; srmG gene; polyketide; tylactone synthesis; antibiotic; tylosin; ss.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  DNA encoding Streptomyces fradiae tylactone synthase domain - for production of tylosin-related polyketide compounds.
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/note= "ORF3 encodes protein shown in AAW22608"
31329. .36071
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/note= "ORF1 encodes protein shown in AAW22606"
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20110. .31284
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P-PSDB; AAW22606, AAW22607, AAW22608, AAW22610.
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This sequence represents the platenolide synthase gene cluster of the invention. This sequence is referred to as the srmG gene, and was isolated from Streptomyces ambofaciens. This sequence encodes the multifunctional proteins which direct the synthesis of the polyketide platenolide. Platenolide is the basic building block of the macrolide antibiotic spiramycin. This sequence was used along with the tylG gene (see AAT80413) to create a hybrid ORF1 sequence (see AAT80415). The tylG gene is the tylactone synthase gene cluster of the invention. The tylG sequence was isolated from Streptomyces fradiae, and encodes multifunctional proteins which direct the synthesis of the polyketide tylosin. The hybrid sequence can be used to transform S. ambofaciens lacking the srmG ORF1 sequence, or S. fradiae lacking the tylG ORF1 sequence, so that they can produce polyketides. The DNA sequence can be modified so as to alter the type of carboxylic acids incorporated and/or the post-condensation reactions performed, thereby resulting in novel tylosin-related
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ------ TCGCACCGTCCT 9075
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16188 CGCCACCTA- 1381 GluGlyThrP 16200 ACAGGACCGC 14101 AlaLeuGlyP 16260 CGCGCACGCCC 1413 16380 GGCGGTGCAG 1416 16490 CCGCCCACAA 1449 GlySex11eT 16490 CCGCCCACAA 1449 GlySex11eT 16490 CCGCCCACAA 1449 GlySex11eT 16599 CCGCCCACAA 1488 16609 GAAGCGACC 1560 GAAGCGACC 1560 GAAGCGACC 1578 AlaArgAlal 16768 CCGTGGCTGC 1506 GLOSETLEUL 16768 GCCCCCCCC 1506 GAAGCGACC 1507 BHISGIAA 1683 -GCGCCCCC 1507 BHISGIAA 16841 GGCCCCCC 1507 GLOSETLEUL 16768 GCCCCCCCC 1507 GLOSETLEUL 16768 GCCCCCCCC 1507 GLOSETLEUL 16768 GCCCCCCCCC 1507 GLOSETLEUL 16768 GCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCC	16188 CGCCACCTA	roProProProProPerargAspLeuThrGlualaTyrLysThrGln 1400	roLeulvsLeulvsProAlaHisGlu	ATGGTGATGGCGCTGCGGAATGG 16319 Db 17302	3	ביים ביישר ביישר נער ביים ניים ביים ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר ביישר 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•	17302	ACCCAACGCGCCACCTCATGCAAACCATGCCCCCGGGACCATGACCACCTCCACACC 17361
_	62	islleproLeuAlaPheAspProThrSerlleProArgGlyIleProLeuAspAlaAlaA 1647
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`	1680	GluAsnArgGlnThrIleIleAsnAspTyrIleThrSerGlnGln 1694
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.	1695	MetHisHis-AsnThrAlaThrAlaMetAlaGlnArgAlaAspMetLeuArgGlyLeuSe 1714
٥	17716	ACCCTCCACCACCACCGCCCACCTACATCGAACTCGGAC17758
پ	1714	rProArgGluSerSerLeuAlaLeuAsnTyrAlaAlaGlyProArgGlyIleIleAspLe 1734
Ω	17759	- CCGACAACACCCTCA 17773
٠.	1734	uSerGlnValProHisLeuProValLeuValProProThrProGlyThrProAlaThrAl 1754
٥	17774	CCACCCTCACCCACACACCCCCCACACCCCCCACCACCAC
>-	1754	aMetAspArgLeuAlaTyrLeuProThrAlaProGlnProPheSerSerArgHisSerSe 1774
۵	17824	CTCACCCCACCCCCACCACCACCCCCAAACCCACCTCCTCACCAACCTCGCC 17874
>-	1774	rSerProLeuSerProGlyGlyProThrHisLeuThrLysProThrThrThrSerSerSe 1794
Ω	17875	AAAACCACCACCACCACCCCCACCACTACACCACCACCACACAACCAAC
>-	1794	rGluArgGluArgAspArgAspArgGluArgAspArgAspArgGluArgGluLysSerIl 1814
۵	17929	-CACACCCACACCCACCTCGAC 17949
>	1814	eLeuThrSerThrThrValGluHisAlaProlleTrpArgProGl 1830
Ω	17950	CTCCCCACCTACCCCTTCCAACACCAGCACTACTGGCTCGAAAGCACACAGCGGGG 18005
>	1830	yThrGluGlnSerSerGlySerSerGlySerGlyGlyGlyGlySerSer 1848
۵	18006	TGCCGGATCCGGTTCCGGTTCCGGGCGGGCAGGACTGCGGGCGG
>-	1848	1848
۵	18066	GGTGGAGTCGCGGTTCTGGGACGCGGTGGCCCGCCAGGACCTGGAAACGGTCGCGACCAC 18125
>-	1849	SerArgProAlaSerHi 1854
۵	18126	GCTCGCCGTGCCCCCCCCCGCCGGCCTGGACACGGTGGTGCCCCGCACTCTCCGCCTGGCA 18185

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'n	885 aThrAlaGluGlyAlaLeuLyBAlaGluLyBLy8	į	: : : :
qq	14116 GCCACCACCTCATCACCCAACGCGCCACCCTCATGCAAACCATGCCCCCGGCACCATG 14175	3 8	
ολ	897 -GluGlyGlySerGlyArgAlaThrThrAlaLysSerSerGlyAlaProGlnAspSerAs 916	3 8	ACTGCGGTCCCC
셤	14176 ACCACCCTCCACACCCCCCCACCACCACCACCACCACCAC	ò	1228 SerIleThr
ò	916 pSerSerAlaThrCysSerAlaAspGluValAspGluAlaGluGlyGlyAspLysAsnAr 936	연	::: 15259 GATGTACGGACGGTCGATGATC
ф	14236 CTCGCCATCGCCGCCATCAACACCCCCCACCTCCTCGTCATCAGCGGCGA 14284	ò	
ò	GLeuLeuSerProArgProSerLeuLeuThrProThrGlyAspProArgAlaAsnAlaSe	q	15319 GGATTCCCGACGAACCGTGGGT
셤	:CCCCACACGTCCAACACATCACCACCCTCTGCCAACAACAA	ò	1231 HisGlyThr
à	rProGlnLysProLeuAspLeuLysGlnLeuLysGlnArgAlaAlaAlaIleProPr	qq	15379 CACGGAACCAGCTACACCCGGA
අු	GGCATCAAAACCAAAAACCTCCCCACCAACCACGCCTTCCACTCCCCCCACACCCC	ò	1234ProA
ð i		qq	15439 GACTICTICGGTATCAGICCGO
g G	ATCCTCAACCAACTCCACCAGCACCCAAACCCTCACCTACCACCCCCACCCCCC	δò	1239
ði í	994 OAlaProProAlaProProProGinAsnbeuGinProGinSerAspAlaProGinGi 1014	අු	15499 CTGGAAACAGCGTGGGAGAGCA
සු .	14449 CICAICACCGCCAACACCCCACCCAACICCICACCCCCACIAALAA 14508	ò	1242 ThrileThrArgileileGly-
ð 1	1014 nProGlySerSerProArgGlyLySeSerProArgGerFroArgaProProArgaBpLySeGrual 1034	qq	15559 ACACCAACGGGGTCTTCGCCG
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ò	1034 aPheAlaAlaGluAlaGluLy8	· qa	15619 GCTCCGGCAGGGTTCGAGGGGT
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ò	1048 sTrpThrSerGlyLeuProPheProValPro	q	15679 CGTGTCGCCTACGCTCTCGGCC
ය		ò	1286 ValThrGlnCysSerLysGluA
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දි ස්	1082 iSProLeuProLeuGlyLeuHisAspinrAlaArgProvalLeuProArgFroProLini 1102	qq	15777 GCGGTCCGGCGAGTGTTCCATG
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1215	ThrargvalProSerAspSerAlaileThrTyrargGly 1227
1228	SerileThr1230 ::: GATGTACGARGGARGATGTCTCTGGCAGGTGGTCAGTGGTGGCCATGACGCGATCGGC 15318
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15319	GGAITCCCGACGAACCGTGGGGGACCTCGACACGCTGTACAACCCGGACCCGGACCAC 15378
1231	HisGlyThr
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15499	CTGGAAACAGCGTGGGAGGCATCGAACACGCCTGCATCAACCCCGACAGCCTCCGTGGC 15558
1242	ThrileThrargileIleGlyGluaspSerProSerArg 1254
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15619	GCTCCGGCAGGGTTCGAGGGGTATCTCGGGCACGGAAGCGCAGGCAG
1267	HisVallleTyrGluGlyLysLysGlyHisValLeuSerTyrGluGlyGlyMetSer 1285
1286	i ValThrGlnCysSerLysGluAspGlyArgSerSerGlyProProHisGluThrAla 1305
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15957	GCTGTCGGACGCGCGCGCGCTCGGTCACCGAATCCTCGCGGTGGTGCGTGGCAGTGCGGT 16016
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1337	CAA1CAGGACGGGGCAAGCAACGGGGCTGACGGGCGCCCAACGGGCCGACCGA
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1342	ProHishisLeuLysGluGlnHisHislleArgGlySerIleThrGlnGlyIlePro 1360
1361	ArgSerTyrValGluAlaGlnGluAspTyrLeuArgArgGluAlaLysLeuLeuLysArg 1380

ysGlyArgileThrargSerMetAlaAsnGluAlaAsnSer	13166 TGGCGCTGCGGAATGGTCTGCTGCCGCGGAction snGluSerSerArgTrpThrGluGluGluMetGluThrAles		647SerGlnCysLysAsnPheTyrPheAsnTyrLysLysArgGlnAs			699 ValValGluAspGluGluMetGluMlaSerGlyValSerGlyAsnGluC 	719 GluGluAlaGluAlaLeuHisAla-SerGlyAsnGluValPro 13516 GCCAGGCCCAGGCCCTGCAGGCCCACCTCACCGACCACCCGGCCTCC	736 BSerGly	751 eProSerProHisThrGlualaalaLysAspThrGlyGlnasr ::: ::: 13636 GCCGACCGCGACCTCCTGCAAGCACTCCAGGCACTCGCCGCAGGCC	769 oProAlaThrLeuGlyAlaAspGlyProProProGlyProPro 		13/9b GCATICALUCGGACAGGGCACCCAACGCCCGGCALGGCCCAAC		80/ -INTRIORIORIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIONIOATAMIOATAMIOATAMIONIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAMIOATAM	826 ulysGluGluGluThrAlaAlaAlaProProValGluGluGlyGluGlu 	846 oAlaAlaGluGluLeuAlaValAspThrGlyLysAlaGluGluPrc	865 uCysThrGluGluAlaGluGluGlyProAlaLysGlyLysAgpAlaGlu
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                                                    The present sequence represents DNA which encodes avermectin aglycon synthase proteins. Also described are: (1) polypeptides encoded by all or part of the DNA; (2) expression vectors containing the DNA; (3) host cells transformed by the vectors; (4) preparation of the polypeptides by culture of the transformants; (5) preparation of avermectin aglycon or its derivatives by culture of transformed avermectin-producing microorganisms; and (6) oligomucleotides of 5-60 bases in length contraining sense or antisense sequences from the avermectin aglycon synthase DNA. The enzymes are useful for the production of modified forms of avermectin and of the intermediates in its biosynthesis, for use as drugs, veterinary drugs and agrochemicals
   2465 nProLeulleMetArgLeuGlnAlaGlyValMetAlaSerProProProProGlyLeuPr 2485
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Avermectin aglycone synthase DNA and proteins encoded by all or part it for the production of avermectin and its derivatives for drug and agrochemical use.
                                                                                                                                                                                                                                                Streptomyces avermitilis; avermectin aglycon synthase; biosynthesis; multifunctional enzyme; polyketide; avermectin; veterinary drug;
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| *tage a
| *Thote = avermectin aglycon synthase protein"
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11857 TTCACAT-----CGGCAACAGAGGCTGAGATTTTCAAATTCATTGACAACGACCTCGGC 11910
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                                                                                                                                                                                                                                                                                                                                                                                  103 eGluSerLysArgProArgLeuGluLeuLeu-ProAspProLeuLeuArg---ProSerP
                                                                                                                                      HighisSerArgAspTyrAlaSerHigLeuSerProGlySerllelleGlnProGlnArg
                                                                                                                                                                                                                    64 ArgArgProSerLeuLeuSerGluPheGlnProGlyAsnGluArgSerGlnGluLeuHis
                                                                                                                                                                                                                                                                                                                                                                                                                        11337 GCCCGCCACCCGCCGCCCCCCCCCCACCACCACCGCGGTGGTGCGGGGACAACGGCGC
                                                                                                                                                                                                                                                                                                   84 LeuligproGluserHisserTyr-LeuProGluLeuGlyLysSerGluMetGluPheIl
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308
1176
1152
152
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Mismatches:
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    30.56%
21.31%
5.17%
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1128 CONTROLCEMENTOATCHCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCCTCC	1989 CGCCGTTGCCGATCACGGGCCCCGGCCACGAATTGCGCGTTGATCGGGCCC 40922 1458 LysTyrAapTh-G1yAlaSerTh-Th-G1ySer_LystySHisAspValAtgSerTeu 476 1457 LieGlySerPCGTGGGCGCCCGGCGCATACGCGCCTCGCCGGCGCCTCGGGG 1477 LieGlySerPCGTGGGCGCCCCGGCGCATACGCGCCTCGCCGGCGCCTCGGGG 1477 LieGlySerPCGTGGGCGCTGCGCGCGCTACGCGCCTCGCCGGCGCCTTGGGG 1477 LieGlySerPCGTGGGCGTGAAACGCCGCGCTGCGCGCGCCTGGGGGCTTGGGGCGCTGGGGCGCTGGGGCGCTGGGGGCGCTGGGGGCGCTGGGGGCGCTGGGGGCGCTGGGGGCGCTGGGGGCGCTGGGGGCGCTGGGGGG	40241 CCGTGCCGCCCGGCTACGCGTTGAGCCCGTTTTGGCCCAGCACCCGCCGG 40182 1727 Y-ProArgGly11e11eAspLeuSerGlNValProHisLeuProValLeuValP 1745 1727 Y-ProArgGly11e11eAspLeuSerGlNValProHisLeuProValLeuValP 1745 1718 CGCCGCGGTTCCGGGGTTGCCGCGTTGCCGCCGTTGCCGCC
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791 gAlaProlleGlubroThrProAlaSerGluAlaThrGlyAlaProThrProProProProAlaserGluAlaThrGlyAlaProThrProProProProProBluI 42691 GGCGCGGCCCGCCGCCCGCCCGCCCGCCGCCGCCGCCGC	SerSerGlyAlaProGln	997 rohlaproProProProGlnAsnLeuGlnBroGluSerApplaBroGlnGlnProGlyS 1017 42197 CGGCCCCCCCCCCAAGA 42168 1017 erSerProArgGlyUysSerArgSerProAlaProProAlaAspLyGGluAlaPheAlaA 1037 1017 erSerProArgGlyUysSerArgSerProAlaProProAlaAspLyGGluAlaPheAlaA 1037 1017 erSerProArgGlyUysSerArgSerProAlaProProAlaAspLyGGluAlaPheAlaA 1037 1037 laGluAlaGlnLysleuProGlyAapProProCysTrpThrSerGlyLeuProPheProV 1057 1057 alProProArgGluVallleLysAlaSerProHisAlaProAspProSerAlaPheSerT 1077 1057 alProProArgGluVallleLysAlaSerProHisAlaProAspProSerAlaPheSerT 1077 1077 yrAlaProProGlyHisProLeuProLeuGlyLeuHisAapThrAlaArgProV 1095 1077 yrAlaProProGlyHisProLeuGlyLeuHisAapThrAlaArgProV 1095 1077 yrAlaProProGlyHisProLeuGlyLeuHisAapThrAlaArgProV 1095 1077 yrAlaProProGlyHisProLeuGlyLeuHisAapThrAlaArgProV 1095 1077 yrAlaProProGlyHisProLeuGly
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Query Match: 5.18* Indels: 652 DB: 4 Gaps: 95 US-09-522-753-5 (1-2517) x AA199683_08 (1-110000) 652 65 US-09-522-753-5 (1-2517) x AA199683_08 (1-110000) 65 65 US-09-522-753-5 (1-2517) x AA199683_08 (1-110000) 65 65 US-09-522-753-5 (1-2517) x AA199683_08 (1-11000) 65 65 US-09-522-753-5 (1-2517) x AA199683_08 (1-11000) 60 60 US-09-522-753-5 (1-2517) x AA199683_08 (1-11000) 65 60 US-09-522-753-5 (1-2517) x AA199683_08 (1-11000) 60 60 US-09-522-753-5 (1-2517) x AA199683_08 (1-11000) 60 60 US-09-522-753-5 (1-2517) x AA199683_08 (1-1000) 60 60 US-09-6000 60 60 60 US-09-6000 60 <t< td=""><td>Db 43365 AAGGAACATTGCCGTTTTGGGGGTTACTTCCGGGAGTTCAGTTCCAGGCCC 43306 Qy 616 ThrGluGluMetGluThAla</td><td>670 LysLeuLysMetGluLysGluArgAsnAlaArgArgLysLysLysAlaProAlaAla 689 43047 CATCTGCGGTC-GAGCGCCGAGCGGTCGCGCGCGCGCGCGCGCCGCGC</td></t<>	Db 43365 AAGGAACATTGCCGTTTTGGGGGTTACTTCCGGGAGTTCAGTTCCAGGCCC 43306 Qy 616 ThrGluGluMetGluThAla	670 LysLeuLysMetGluLysGluArgAsnAlaArgArgLysLysLysAlaProAlaAla 689 43047 CATCTGCGGTC-GAGCGCCGAGCGGTCGCGCGCGCGCGCGCGCCGCGC

1113	GGCGGAGTGACCTCGGCGACGACTTCTGGGATCTCGTCCTCCGAGCAGCACGCGATC 20106 1TrpGludspargProSerSeralaGly	from base 800001 (My 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by 10000 by	Length: 110000 Matches: 530 Conservative: 186 Mismatches: 852
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(Mycobacterium tuberculosis strain H Accession Aai99683 110000 530 7e: 186 852

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1122 leglyA	λο —	ACCCACCCCGTCTTCGCCGCCGCCACTCTGCACCCCCCCC	g (
 14845 TCACCG	අු		ð 1
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	2 qa	OProAlaThrLeuGlyAlaAspGlyProProProGlyProPro	ò
	S &	13636 GCCGACCGCGACACTTCCTGCAAGCACTCCAGCACTCGCCGCAGGCGAACCCACCC	qq
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14617 GACAAC	음 &	 13576 GTCGGGTACACCCTCGCCCACGCCGTGTTCGACCACCGCGCCACCCTCATCGCC 13635	qq
1048 sTrpTh	ò	8SerGlyProAlaThrValAsnAsnSerSerAspThrGluSerIl	ò
14557 CACGGC	qu	13 GIUGIUATAGIUATAGAGUTAGAGAGAGAGAGAGAGAGAGAGAGAGAGAGA	∂ €
1034 aPheAl	è	CTGGTGTCGGCCAAGTCGCAGCCGGCCCTGCGC	a
: 14509 GCCCGC	qu	ValValGluAspGluGluMetGluAlaSerGlyValSerGlyAsnGluGluGluMetVal	ò
1014 nProGly	& 	13423 GACGAGGATGCCGGTAGTGGCGATGAGGCTGCTGCGGCAGTCCAGGGGTGTGGCCGTGG 13482	q
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975 ollegl	ò	13303	B &
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	q a ,	13166 TGGCGCTGCGGAATGGTCTGCCGCGGGACGTTGCATGTGG 13207	අ
897 -GluGly	ò	GluGluAlaIleThrProGlnGlnSerAlaGluLeuAlaSerMetGluLeuA	ò
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865 uCysŢḥ	ò		•

1194		3
1180	62 AlaProPheSerGlyValLysGlnGluGlnLeuSerProArgGlyGlnAlaGlyPro 1 	116
15024	### Bandarion from the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of the control of th	1496
14964	05 GTGGCCAAGTCTCCGCCAGGAGGAGGACGAAGGCTGTTGCGCCCTGGTGCGCACCCATGCG	1490
1141	22 leGlyAlarleSerGlnGlyMetSerValGlnLeuHisValProTyrSerGlu-HisAla	113
1122		111
	:: 88 TACCCCTTCCAACACCACTACTGGCTCCAACCACCGGCAAGCCGAGGGACCCG	1478
1117	02 leSerAsnProProProLeulleSerSerAlaLy8Hi8ProSerV	110
1102	82 isProLeuProLeuGlyLeuHisAspThrAlaArgProValLeuProArgProProThr1 3	108
1082 14736	63 eLysalaSer-ProHisalaProAspProSerAlaPheSerTyrAlaProProGlyH	106
14676	17 GACAACCTCCCAACACCCCCACCACCACCTCACCCTCACCCCCC	1461
1063	48 sTrpThrSerGlyLeuProPheProValProProArgGluValll	104
α	34 aPheAlaAlaGluAlaGlnLysLeuProGlyAspProProCy 3	-
1034 14556	14 nProGlySerSerProArgGlyLysSerArgSerProAlaProProAlaAspLysGluAl 1 	101
14508	-::	1444
4	94 oAlaProProAlaProProProProGlnAsnLeuGlnProGluSerAspAlaProGlnGl	
994	75 olleginValThrLysValHisGluProProArgGluAspAlaAlaProThrLysPr 5	97.
14388	29 GGGATCAAAACCAAAAACCTCCCCACCAACAACGCCTTCCACTCCCCCCAAAACCAACC	1432
	56 rProGlnLvsProLeuAspLeuLvsGlnLeuLvsGlnArgAlaAlaAlaIleProPr	
956 14328	36 gLeuLeuSerProArgProSerLeuLeuThrProThrGlyAspProArgAlaAsnAlaSe 5	93
14284		1423
	SorCorb aThrOsSorb sections SorCorb	5
916 14235	97 -GluGlyGlySerGlyArgAlaThrThrAlaLyBSerSerGlyAlaProGlnAgpSerAg 5 	1417
14175	16 GCCACCACCTCATCACCCAACGCGCCACCTCATGCAAACCATGCCCCCCGGCACCATG	1411
968	35 aThrAlaGluGlyAlaLeuLysAlaGluLysLys	88
14115	os ucystinatudidadatudidatus yriohidatay by braphagatuda da	1405
882	55 uCvsThrGluGluAlaGluGluGluGlvProAlaLysGlyLysAspAlaGluAlaAlaGluAl	8

11953 -GGCCCTGCTAGGACGTGAATGGCGAATGAAGCGAAGCTCCTGGAATACCTCAA 12011 334 uLysGlnPheProGluIleArgLysGlnArgGluLeuGlnGluArgMe 350 ::::	365 aAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSerGluGlnGlu 383	400 rAspalaAspGlnGlnArgIleLysPheIleAsnMetAsnGlyLeuWetAlaAs 400 rAspalaAspGlnGlnArgIleLysPheIleAsnMetAsnGlyLeuWetAlaAs 1219 cGACCCGGACCGCTCAGGAACCAGTTACACGCGCAGCGGAGGGTTTCTCTATGA 418 p	12279 GGGGGGGACTTCGACGGTTCTTCGAGTTGTCACGGGGGGGG	447 12399 CCCGAGGTCCATGCGCGGACCGGGGTTTTCGTGGGGATCATCCGCGAGGACTA 12458	461 12518	12578 12578 484	494 12698 495	12758 496 12818	y 509 nGlnProMetProArgSerSerGlnGluGluLy8AspGluLy8GluLy8GluLy8GluLJ 529 ::::::::::::::::::::::::::::::::::::	12938 TCATCCGCCAGGCTTTGGCCAACGCACCTCTCCCCTGCCGATGTGG 543 luaspleuleulysGlulysThraspaspThrserGlyGluaspasnaspGlulysGlua 12986 ATGCGGTGGACGGCCACGGGACGCACCTGGGGACCCGATCGAGGCTCAAGGCC
8 6 8 6 8	2 8 2 6	6 6 6	a & a	6666	3 8 8 8	3 8 8 8	8 8 8	3 8 8 8	% d	3, A & A
Alignment Scores: 3.63e-12 Length: 30690 Score: 685.00 Matches: 704 Percent Similarity: 30.18\$ Conservative: 300 Best Local Similarity: 21.16\$ Mismatches: 1193 Query Match: 4.18\$ Gaps: 148	US-09-522-753-5 (1-2517) x AAH79277 (1-30690) Qy	11235 84 11286	Qy 103 ecinbelly had greater the prediction of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of the property of	142 hrGlyLysLeuGluProvalSerProProSerProProHisThrAspProGluLeuGluL 		Qy 202 GluGluAlaAlaLysProProGluProGluLysProValSerProProProlle 219 :::	Db 11654 ACCTCCACACACTCCAGCCACAACGGGACAGGTGT 11693 Qy 239 UAlaAlaHisArgIleLeuGlUGlyLeuGlyProGlnValGluLeuProLeuTyrAs 258 Db 11694CGCCCCGGGAGGTCGACAAACTCGAATCCGCCCTCTC 11738	Qy 258 nGlnProSerAspThrArgGln	Db	QY 299 -CysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGluArgIleGluAs 318

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----GCCCGTTGGC 36086
                                                                                                     36085 GCCGTTGCCGATCAACGGGCGCCCCGTGAGCGCGACACTTTGCGCATTGACGCGGCCCAG 36026
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           The present invention relates to the production of modified derivatives of a vermectin aglycone synthase (AAS) derived from Streptomyces avermitilis. The activity of an acyl carrier protein (ACP), beta-ketoacyl carrier protein synthase (KS), acyltransferase (AT), beta-ketoacyl carrier protein reductase (KR), acyltransferase (DH), beta-ketoacyl carrier protein reductase (KR), dehydratase (DH), enoyl reductase (ER) and/or thoseterase (TE) domain may be reduced or suppressed. The process can be used in the production of drugs, vetinary drugs and pesticides. The present sequence is a fragment of the S. avermitilis genome
            2426 gProProSerValSerSerValHisSerGluGlyAspCysAsnArgArgThrProLeuTh
                                                                        2446 rAsnArgValTrpGluAspArgProSerSerAlaGlySerThrProPhe---ProTyrAs
                                                                                                                                   nProLeuIleMetArgLeuGlnAlaGlyValMetAlaSerProProProProGlyLeuPr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    New modified avermectin aglycone synthase derived from Streptomyces avermectilis used in production of 22,23-dihydroavermectin Bla used
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Sequence 30690 BP; 5355 A; 12455 C; 8617 G; 4263 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                  Avermectin aglycone synthase; AAS; avermectin derivative;
drug production; vetinary drug; pesticide; ds.
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                                           Example 2; Page 58-123; 257pp; Japanese.
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11971. .30690
/*tag= b
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P-PSDB; AAG65264, AAG65265.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Streptomyces avermitilis.
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drugs and pesticides
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·	546 41474 566	GGTGACGGAATCICGAAGATGCGICGAITCGIGGCGAAGCGGAAGGGAAG	0y 6124 AAGGACATTCGGCGGCTGTTCTCGGCGGTTGCACCTGGCGCTTGGCGCGTGGCTGGC
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Reinhard C, Randazzo F, Kennedy GC, Pot D, Lamson
Crkenjakov R, Drmanac S, Dickson M, Labat I;
z D, Kita D, Garcia V, Jones LW, Strache-Crain B;
                                                                                                                                             Human, cytostatic, gene therapy; colon cancer, prostate cancer;
breast cancer; lung cancer; cancer detection; ss.
                                                                                                                                                                                                                                                                                                                                                                                                  Library of polynucleotides for diagnosing a cancerous state of mammalian cell and detecting cancer, particularly of the colon prostate, comprises 3351 human polynucleotide sequences.
                                                                                                                        Novel human polynucleotide, SEQ ID NO: 2976.
                                                                                                                                                                                                                                                                                                                                                                                                                                             Claim 9; Page 988; 1046pp; English.
------HisAlaTrpAsp 2499
                    8763
                                                            BP.
                   8782 AGCGGCCATCGGGGTGGGAT
                                                                                                                                                                                                                                         30-JUN-2000; 2000WO-US018374.
                                                                                                                                                                                                                                                            99US-0142310P.
                                                            AAF67220 standard; cDNA; 427
                                                                                                     (first entry)
                                                                                                                                                                                                                                                                                                                                      Kassam A, Reinhard C,
Drmanac R, Crkenjakov R
Leshkowitiz D, Kita D,
                                                                                                                                                                                                                                                                                            (CHIR ) CHIRON CORP. (HYSE-) HYSEQ INC.
                                                                                                                                                                                                                                                                                                                                                                               WPI; 2001-091805/10.
                                                                                                                                                                                                WO200102568-A2
                                                                                                                                                                                                                                                              02-JUL-1999;
02-JUL-1999;
                                                                                                                                                                              Homo sapiens.
                                                                                                                                                                                                                                                                                                                             Williams LT,
                                                                                                     09-APR-2001
                                                                                                                                                                                                                     11-JAN-2001
2496
                                                                                 AAF67220;
                                         RESULT 27
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The present sequence is one of 1351 sequences in a library of human polynucleotides. The library is used to detect differentially expressed genes correlated with a cancerous state of a mammalian cell and can detect colon, prostate, breast and lung cancer. The library can be used to produce probes for detection of mRNA and to produce additional copies of the polynucleotides. The probes can be used for chromosome mapping of the polynucleotides of detection of transcription levels. Ribozymes or antisense oligonucleotides can be generated. The polynucleotides and their gene products are used as genetic or biochemical markers (e.g. in blood or tissues) that will detect the earliest changes along the carinogenesis pathway and/or monitor the efficacy of therapies and products are used in pharmaceutical compositions to the cancers and produce against them can be used in pharmaceutical compositions to the cancers and brackles golders such as neoplasia, hyperplasia dysplasia and

Sequence 427 BP; 94 A; 143 C; 159 G; 31 T; 0 U; 0 Other;

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427
135
0
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0
      Length:
Matches:
Conservative:
Mismatches:
Indels:
      1.03e-13
689.00
97.83%
97.83%
                       Percent Similarity:
Best Local Similarity:
Alignment Scores:
                                      Query Match:
DB:
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US-09-522-753-5 (1-2517) x AAF67220 (1-427)

AlaProlleGluProThrProAlaSerGluAlaThrGlyAlaProThrProProAla 811 792

4 5 4 5 4 5 4 5 6	8 8 8 8 8	8 8 8 8 8 8	8 8 8 8 8 8	8 8 8 8 8 8
ACGGCGCAGCACCACCACCACCACCACCACCACCACCACC	11172 ACACCGGACGA			1805 pargAspArgGluLysSerIleLeuThrSerThrThrThtValGluHisAlaPr
8 2 8 2 8 2 8 2	8 6 8 6 8 6 8	8 6 8 6 8 6 8	6 6 6 6 6 6 6	\$ 8 \$ 8 \$ 8 \$

10472 CA	CAGCAACACCCGACGCCACCGAACCC 10447
1885 lyIle	IlelleThralaValGluProSerLysProThrValLeuArgSerThrSerThrSerS 1905
n o	ovalArgProAlaAlaThrPhePr
1925 eu 10358 CC	eu-aspGlyValTyrProThrLeuMetGluProValLeuLeuProLysGluAlaProArg 1944
1945 Va 10298 CA	ValalaargProGluargProArgAlaaspThrGlyHisAlaPheLeuAlaLyg 1962
63	ProproAlaArgSerGlyLeuGluProAlaSerSerProSerLysGlySerGluProArg 1982 ::
1983 Pr 10190 CC	Pro
1988 Va 10130 CC	ValSerGlyHisAlaThrIleAlaArgThrProAlaLy8AsnLeuAlaBroHisHisAla 2007
98	SerProAspProProAlaProProAlaSerAlaSerAspProHisArgGluLys 2025
9 6	ThrGlnSerLysProPheSerIleGlnGluLeuGluLeuArgSerLeuGlyTyrHisGly 2045
9 6	SerSerTyrSerProGluGlyValGluProValSerProValSerSerProSerLeuThr 2065 :::
929	HisaspLysGlyLeuProLysHisLeuGlu-GluLeu
078	
	2101
01	1 8
2115 rf 9714 CC	srSerSerProLeuLeuGlnThrAlaProGlyValLy8GlyHisGlnArgValValThrLe 2135 :::::
2135 w	uAlaGlnHisIleSerGluValIleThrGlnAspTyrThrArgHisHisProGlnGlnLe 2155 ::::::: ::::::::
2 2	erAlaProleuProAlaProL AGAACCCGCACCACCGCCAA
175	pleuargargproproseraspleuTyrLeuProProProAspHisGlyAlaProAlaar 2195 :: :: ::

Oy 1278 uSerTyrGluGlyGlyMetSerValThrGlnCysSerLysGluAspGl 1294	1314 tGluGlyArgValGlyArgAla1leSerSerAlaSerIleGluGlyLeuMetGlyArgAl 	1334 alleProProGluArgHisSerProHisHisLeuLysGluGlnHisHisIleArgGlySe 13138 TGGTCCAGCATCCGGGCATTCG	luAspTyrLeuArgArgGl	1374 ualaLysLeuLeuLysAxgGluGlyThrProProPro	Qy 1389 oSerArgAspLeuThrGluAlaTyrLysThrGlnAlaLeuGlyProLeuLysLeuLysPr 1409 :::	Oy 1409 oAlaHisGluGlyLeuValAlaThrValLyBGluAlaGlyArgSer1leHisGlu 1427	Qy 1428 -IleProArgGluGluLeuArgHisThrProGluLeuProLeuAlaBroArgBroLeuLy 1447 ::: ::	Oy 1447 sGluGlySerlleThrGlnGlyThrProLeu-LysTyrAspThrGlyAlaSerThrThrG 1467 : : : : : : : : : : : : : : : : : : :	Qy 1467 lyserLysHisAspValArg	Qy 1477 leGlySerProGlyArgThrPheProProValHisProLeuAspValMetAlaAspAlaA 1497	Qy 1497 rgAlaLeuGluArgAlaCysTyrGluGluSerLeuLysSerArgProGly 1513	Oy 1513 1513 1513 1513 1513 1513 1513 1513	1514ThralaSerSerSerGlyGlySerIleAlaArgGly-	Db 11631 CAGGTGATCCGATACCGCACGCCACCCCACGGGGGGGGGG	11571 GGGTTGGTGGCGGGGTTGGTGGGAGTCGGTGCCCAGTAGTGGCGGCGTTGGAACGGATAC	<pre>Qy 1532GluLeuGlyLysProArgGlnSerProLeuThrTyrGluAspHisGlyAlaProP 1550</pre>	1550 heAlaGlyHisLeuProArgGlySerProValThrMetArgGluProThrPro-	Db 11460 TCAACACTCATCCGGCGACGAAGGCCTCACCGACGACGTCAAAAAACGGCCCCAC 11404 Qy 1567
1029 ProAlaAspLysGluAlaPheAla	13241 TCAGACCGGTCGTGTTGAGGTGGTTGCGCAGTTCGAGCGCGTGAGCGAGTCGAAGC 13182 1042	1054 ProPheProValProProArgGluValIleLysAlaSerPro-HisAlaProAspProSe 1073	oProGlyHisPro TCCCGCCACCCG		1106ProproLeulleSerSerAlaLys			1139 uHisAlaLysAlaProValGlyPro		luse		12618 TGCTGGGTGAGTTCGTCGAGCAGCAGCAGCAGTCTTCGGACGGA	12558 ATGTGTCCGGCGGCGAGCGTGTCGATCACACCGTCCAGGACTCCGGCGGTGTGG 12502		1225 TyrArgGlySerIleThrHisGlyThrProAlaAspValLeuTyrLysGlyThrIleThr 1244 ::: ::: ::::	1245 ArgilelleGlyGluAspSerProSerArgLeuAspArgGly 1258		

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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     The invention relates to genes and proteins involved in the biosynthesis of macrolides by microorganisms. In particular it relates to the nucleic acids forming the biosynthetic locus for rosaramicin (a 16-member macrolide artibiotic) from Micromonospora carbonacea. The invention is useful for the biosynthesis of macrolides by microorganisms. It allows direct manipulation of macrolides and related chemical structures by chemical engineering of the proteins involved in the biosynthesis of rosaramicin. It is useful to catalyse certain biochemical reactions, in withoo or in vivo, to direct or enhance the synthesis or modification of a polyketide substrate or its precursor. The present sequence is M. carbonacea polyketide synthase (PKS) type I gene cluster. (Updated
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ArgAlaThrGluProArgTyrProProHisSerLeuSerTyrProValGlnIleAlaArg 32
                                                                                                                                                                                                                                                                                                                                                                                                               Novel isolated polypeptide involved in biosynthesis of macrolides by microorganisms, useful for biosynthesis of macrolides by microorganisms, preferably for biosynthesis of rosaramicin.
                                                                                                                                                                                                                                                                                                                                                                 P-PSDB; AAE36995, AAE36996, AAE36997, AAE36998, AAE36999, AAE37000,
AAE37001, AAE37002, AAE37003, AAE37004, AAE37005, AAE37007,
AAE37008, AAE37009, AAE37010, AAE37011, AAE37012, AAE37013.
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1027
                                                                                                                     /product= "Polyketide synthase #18"
/note= "CDS does not include start
                                                                                 /product= "Polyketide synthase #17"
57657. .59123
                                           'product= "Polyketide synthase #16"
            "Polyketide synthase #15'
                                                                                                                                                                                 synthase #19'
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Matches:
Conservative:
Mismatches:
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/product= "Polyketide
                                                                                                                                                                                                                                                                                                                                                                                                                                                                Example 2; Page 59-94; 206pp; English.
                                                                                                                                                                                                                                                                                                        (ECOP-) ECOPIA BIOSCIENCES INC.
            /product= "Po]
54897. .56342
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/*tag= 8
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Best Local Similarity:
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     189 lnIleSerLysLeuLysLysLysGlnGlnGlnGlnGluGluAlaAlaLysProProG 209
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---ACAAACCACCCAACAACCCCGACAGCATCCCCTCCACCGC
                                 ---GCCCCACCCACCATCATCACGCGCAGCGACCCACCACCACCACGGCGCACATCCGG
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53 LeuSerProGlySerIleIle------GlnProGlnArgArgArgProSerLeuLeu
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                                                                                                                  70 SerGluPheGlnProGlyAsnGluArgSerGlnGluLeuHisLeuArg-ProGluSerHi
                                                                                                                                                                                                                                                                                            15856 CTCCACAAACCCCGAA--
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ThrHisThrAspValGlyLeuLeuGluTyrGlnHisHisSerArgAspTyrAlaSerHis

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CAGGTCAACCAACCGCGCGCCCCCCCCGCGAACACACCCCTTCCAGTC

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product= "Polyketide synthase #13"
trans1 except= (pos:51221. .51223, aa:Xaa)
note= "Xaa corresponds to amino acids from position 452-
                                                                                                                                                                                                                                                                                                                          Micromonospora carbonacea polyketide synthase (PKS) type I gene cluster.
                                                                                                                                                                                                                                                                                                                                                                                        Macrolide; rosaramicin; polyketide; polyketide synthase; PKS; enzyme;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  product= "Polyketide synthase #10"
note= "CDS does not include start codon"
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note= "CDS does not include start codon"
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/note= "CDS does not include start codon"
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9868. .51226
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note= "CDS does not include start
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     product= "Polyketide synthase #9"
6400. .47794
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product= "Polyketide synthase #3"
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9144. .24921
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/product= "Polyketide synthase #2"
complement(2629. .3861)
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            *tag= g
product= "Polyketide synthase #7"
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      complement (1728. .2522)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Location/Qualifiers
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3081 AGCGGCCATCGGGGTGGGAT 3062
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   3640 -----CGAACACATACGAAACACGACCGAC 3615
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                2303 -GluTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetProAlaIleThrGlyTh 2322
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  2322 rGlyLeuMetThrTyrArgSerGlnAlaValGlnGluHisAlaSerThrAsnMetGlyLe 2342
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            | :::|||:::||| | | 3554 AACCGCGCCATACACCCGCCATACACCGCGCAAACCC 3498
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      2382 olleThrAlaAlaAspGlyArgSerAspHisThrLeuThrSerProGlyGlyGlyGlyLy 2402
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              3497 CGCAAACCCAACGGAT---CAATCCCCGCACGCTCCAACGCCTCCC----- 3455
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        3405 ACGCGGCGÁCACACCÁAAAAACTCCGCATCGAACATCCCCGCGTCATACACAAACCCCCC 3346
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                                                                                            2203 yLysArgSerProGluProAsnLysThrSerValLeuGlyGlyGlyGluAspGlyIleGl 2223
                                                                                                                                                                                                                 2223 uProValSerProProGluGlyMetThrGluProGlyHisSerArgSerAlaValTyrPr 2243
                                                                                                                                                                                                                                                             2263 yAsnThrSerGlnProProAlaPhePheSerLysLeuThrGluSerAsnSerAlaMetVa 2283
                                                                                                                                                                                                                                                                                                                                       2243 oLeuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlySerLysSerProGl 2263
                                                                                                                                                                                                                                                                                                                                                                                              T------GACACGCCAAATGCAACGCCACCAACGACGACGAACACGCCGTATCAACC 3663
                                     3860 greceardedadeaceacadaaacececaacacecearcaceceaaacecaceaceace
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                                                                                                                                                             3800 CGAGAAACTCCACAAACAACCCCGGCG---
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SEPROVALSERSERPROSERLEUThr 2065 aralaSerAspProHisArgGluLys 2025 ubeuArgSerLeuGlyTyrHigGly 2045 31yGluLeuArgProLysGln---- 2090 :::||||||::|||| ccgcccaTAcGACcCaACACGCT 4121 |||::::::||| cccargcgccrc-----ca 4073 --- ProLeuProGluSerGlnProSe 2115 YrThrArgHisHisProGlnGlnLe 2155 CATTCGACCCACCATCCTGATTCACC 3957 roGlyAlaSerCysProValLeuAs 2175 roproAspHisGlyAlaProAlaAr 2195 ThrHisCysProLeuGlyGlyThrL 1925 ACG-----ACCG 4658 spThrGlyHisAlaPheLeuAlaLys 1962 srProSerLysGlySerGluProArg 1982 GGATCGTCGAATTCGGGGGCCTGCT 4430 aLysAsnLeuAlaProHisHisAla 2007 AACGCCGACACCCCGCACGACGCA 4370 roValLysieuGlyGlyGluAlaAl 2101 CCACCAACGCCCCCAATCACCCGA 4014 ysGlyHisGlnArgValValThrLe 2135 cceargaccaccrccccc---- 3911 -----SerGluGlyGl 2203 III::: III 1 LeuLeuProLysGluAlaProArg 1944 -------LeuValProPro 1987 ren----- 2011 STGCGTATGACCAATATTCGACTTCA 4181 cacccccacaccrccccccaccc 3861 ValLeuArgSerThrSerThrSerS 1905 GACCAACCCÁCATTCACCAGACCGG 4598 - Accedecegearaagegececaaag 4541 2195 gGlySerProHis-------

6563 TTCCCGCGAGGGCGCCATTCCCGG 1294 YAXGSerSerSerGlyProProHi	6503 TAGATCCCCGAGCCGCGTACCGCG	6457CGCCGAGGCCT	1334 alleProProGluArgHisSerPr	6437 IGGTCCAGCATCCGGGCATTCG	1354 rlleThrGlnGlylleProArgSe :: 6409ACGGAAGGT		6400	1389 oSerArgAspLeuThrGluAlaTy	6356 CGCCCGAACGCCCACACTGCGCA	6320 GGGCCTCCTGGGGCACCACAGCG	1428 -IleProArgGluGluLeuArgHi	::: 6260 GCTCCCGCGTCACCGATGGCC	1447 sGluGlySerIleThrGlnGlyTh	6203 ATGGAGGGTGTTGAGGATGCGCG	1467 lySerLysLysHisAspValArg-	6162AGGAGACACGCCCCCGAC	14// leclyserrioclysiginization //ll	1497 rdala[]undrdalaCvsTvrC	6047 CTGCGAATGAGCGCTGCTGTG	1513	5990 TCGGCGATGAGTAGCCAGCGGTTG	1514	5930 CAGGIGATCCGATACCGCACGCCA	1526	5870 GGGIIGGIGGCGGGGIIGGIGGGA	1532GluLeuGlyLysProArgG	5810 GTCGGCAGGTCAACCAACCGCGCG	1550 neAlaGIYHIBLGUPrOArgGIYS		5702 CCACCATCATCACGCGCGCAGCGAC
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ą	5930 C	AGGIGATCCGATACCGCACGCCACCGGGTCGGCGGTGGTGGTGTCGCCCGTGGGG 5871
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ą	5870 G	GGGTTGGTGGCGGGGTTGGTGGGAGTCGGTGCCCAGTAGTGGCGGCGTTGGAACGGATAC 5811
٨	1532	GluLeuGlyLysProArgGlnSerProLeuThrTyrGluAspHisGlyAlaProP 1550
ą	5810 G	TCGGCAGGTCAACCAACCGCGCGCCCGCCCCCGCGAACACACCCTTCCAG 5760
≿ .	1550 he	heAlaGlyHisLeuProArgGlySerProValThrMetArgGluProThrPro- 1567
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          Novel isolated polypeptide involved in biosynthesis of macrolides by microorganisms, useful for biosynthesis of macrolides by microorganisms, preferably for biosynthesis of rosaramicin.
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ACTCATCCCGCCGACGAACGCTCACCCACCGACGTCAGAAACCG-------
                                                                                                                                     rosaramicin; polyketide; polyketide synthase; PKS; enzyme;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 BP; 1361 A; 3414 C; 5846 G; 2795 T; 0 U; 0 Other;
                                                                                                Micromonospora carbonacea polyketide synthase (PKS) type I gene
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Matches:
Conservative:
Mismatches:
Indels:
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Claim 1; Page 120-128; 206pp; English
                                                                                                                                                                                                                                  Location/Qualifiers
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P-PSDB; AAE36999.
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Best Local Similarity:
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07-AUG-2003
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                                                                                                                                       Macrolide;
    AAD55815;
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ValSerGlyArgProSerSerArgLysAla-LysSerPro---Ala 2418 rGlyAspArgProProSerValSerSerValHisSerGluGlyAsp 2438 rProLeuThrAsnArgValTrpGluAspArgProSerSerAlaGly 2458 oTyrAsnProLeulleMetArgLeuGlnAlaGlyValMetAlaSer 2478 2244 7135 LysAlaLeuMetGlyLysTyrAspGlnTrpGluGluSerProProL 2363 AlaAlaAspGlyArgSerAspHisThrLeuThrSerProGlyGlyG 2400 ACCACGTCCG----ACCACC 6851 yLeuProAlaGlySerGlyProLeuAlaGlyProHisHisAlaTrp 2498 cecescearecreatececerececececeaecreceaecece crcceaggccrccaccarcarcarccccccggcrcgcaccr 7395 lyGluGlnThrGluProSerArgMetGlySerLysSerProGlyAs 2264 TCGGGCGCACA-----CTAGCTTCGCTGCGTCCGCAAGG 7347 cacatececaececteaectetetatesagececaecaecaeca 7287 GlnGlulleAsnLysLysLeuAsnThrHis---AsnArgAsnGluP 2302 -----G 2303 CTCCGTCCGCTCCAGCAACGCTCCTTCCTCGCTGCCCGCGGCCGCG 7167 GlnProGlyThrGluIlePheAsnMetProAlaIleThrGlyThrG 2323 ArgSerGlnAlaValGlnGluHisAlaSerThrAsnMetGlyLeuG 2343 GCCCGCGACT------CGAAGTGCGTCCA 6881 ro---AlaPhePheSerLysLeuThrGluSerAsnSerAlaMetVa 2283 ::: ||| :::| ccacgactcccactgacgtacagcgccacttccagcgcaaacagc 7227 AAAGCC-----TCTTCCCATCC 7057 roAsnLysThrSerValLeuGlyGlyGlyGluAspGlyIleGluPr luGlyMetThrGluProGlyHisSerArgSerAlaValTyrProLe caaccacgercgagara---SCTCGTCTC 6782 sProLeuLeu 2506 13416 BP

1065 AlaSerProHisAlaPro 1156 CGCTCGCCCAAGCACCC 1082 HisProLeubroLeuGl) 11506 GCACCACGTCGC 1102 IleSerAsnProProPro 111461 TCGCCCACGTCGCCCCCGCCCCCCCCCCCCCCCCCCCCC	1305 aAlaProLysArgThrTT
8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6 8 6	6 6 6 6 6
12646 CCTGGGCCCAGTCCACCTGGTGCACCTGCACTGTGCGCACTGCCCCGTCG 823	
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------ACCACCCCACTCCCCGTGCTCTTCCAGCCACT 11462 TCGGCGCGTGATC-----GCGCTGCTGCGG 10872 ::||| ::: ccgrgccgcgcaccacacccagacccgcrrccccrgcrgcrgc 10812 |||::: |TCAGCACCAGCACCCCGACTCCCTCGCCACGGCCATAACCGTCC 10752 CTCGCAATCGCCCCGGCGCAGCGCTCGCAGGCCAGGTG---- 10600 || ::: ::: CT-----GCGCATTCACCGCGCCTCGTCTCGACCCGACAGCA 11408 ||| GCGCGGCGAGCTGTGCAGCGTCGCCGGCAGCGACGGA 11153 hrHis---GlyThrProAlaAspVal-LeuTyrLysGlyThr-- 1242 |||||||:: caacaccaacacaaaracarcaaracaacaccaccaccars 11507 GCAC-ACCCCTGCGATGCCCGCGGGGACTCGAGATGACCGATA 11094 InGlyMetSerValGlnLeuHisValProTyrSerGluHisAla 1141 erileThrLysGlyIleProSerThr------ 1215 ysGlyHisValileTyrGluGlyLysLysGlyHisValLeuSer 1279 ervalThrGln-----CysSer 1290 SerSerSerGlyPro------ProHisGluThrAl 1305 TyrAspMetMetGluGlyArgValGlyArgAlaileSerSerAl 1325 IleArgGlySerIleThrGlnGlyIleProArgSerTyrValGl 1365 lyLeuHisAspThrAlaArgProValLeuProArgProThr 1101 roLeulleSerSerAlaLysHisProSerValLeuGluArgGln 1121 rovalThrMet----- 1150 alLysGlnGluGlnLeuSerProArgGlyGlnAlaGlyProPro 1181 roThralaGlnGlualaSerValLeuArgGlyThrAlaLeuGly 1201 roSerAspSerAla------IleThr 1224 lyGluAspSerProSerArgLeuAspArgGlyArgGluAsp--- 1261 coAspPro-----SerAlaPheSerTyrAlaProProGly 1081

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14757 GCGATGCGCCGCGAACGCGTCGAGGAAGCTGTTGGCCGCGCTAGTTGCTCTGGCCCGC 14698
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5323 ACGG---CGCGCGCGCTCGGGTAGTCGAACACCAGCGTCGCCGGGAGGGGATCTCGGCT 15267
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               4877 CACAAGTCCGGCGCGCTGGATACGTGAGACTCCGCCTCGCCCAGGTGCGCTGTCATGCC 14818
                                                                                                                                                               299 CysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGluArgIleGluAsn 318
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            249 GlyProGlnValGluLeuProLeuTyrAsnGlnProSerAspThrArgGlnTyrHisGlu 268
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               269 Asnile-----LysileAsnGlnAlaMetArgLysLysLeuileLeuTyrPheLys 285
                                                                                                                                                                                                                                                                                                                                116 oLeuLeuArgProSerProLeuLeuAlaThrGlyGlnProAlaGlySerGluAspLeuTh 136
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-LeuSerProGlySerileIleGlnPro
                                                                                              15620 ccrecérceaagacarceararcegeagaarceacéregegeaggageere
                                                                                                                             73 nProGlyAsnGluArgSerGlnGluLeuHisLeuArg-----ProGluSerHisSe
                                                                                                                                                                                                                              15500 ATGGCGTCCTTCCCTTCCGCCAGGAGACGCCAGTAGTCTTCCGGCGTCGCCACGCCCCT
                                                                                                                                                                                                                                                                ----PhelleGluSerLysArgProArgLeuGluLeuLeuProAspPr
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                                                               ----ProSerLeuLeuSerGluPheGl
                                                                                                                                                                                                 90 rTyrLeuProGluLeuGlyLysSerGluMetGlu----
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                                                               62 Gln-ArgArgArg
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                New recombinant DNA vector for preparing hybrid polyketide synthase enzymes comprises all or a portion of one or more genes in the tmbA gene cluster of Sorangium cellulosum.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              The invention describes a recombinant DNA vector that comprises all or a portion of one or more genes in the tubA gene cluster. Also disclosed is the host cell that comprises the tubA genes of Sorangium cellulosum or fragments of those genes. The DNA vector is useful in preparing bybrid polyketide synthase (PKS) enzymes and the polyketides produced by such hybrid enzymes. The gene products of the tubA cluster can be used to synthesize the polyketide tombamycin. This sequence represents the Sorangium cellulosum tubA gene cluster tubB open reading frame.
                                                                                                                                                               recombinant DNA vector; tmbA gene cluster; tmbA; tmbB; tmbC;
polyketide synthase enzyme; PKS enzyme; hybrid PKS enzyme; tombamycin;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GlubroArgTyrProProHisSerLeuSerTyrProValGlnIleAlaArgThrHisThr
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Matches:
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Mismatches:
Indels:
                                                                                                                                  Sorangium cellulosum tmbA gene cluster tmbB DNA.
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1. .20922
                                  ADC26981 standard; DNA; 20922 BP
                                                                                                                                                                                                                                                                                                                  "tmbB"
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31-AUG-1998; 98US-00144085.
15-FEB-2001; 2001US-0271245P.
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21428 AGGCCTCGATCACGTCTTCCCCTCGGCCAAGAGACGCCAGTAGTCTTCCGGCGTCTCG 21369 2216	2258 ySerLysSerProGlyAsnThrSerGlnProProAlaPhePheSerLysLeuThrGluSe 21190 ACGTCTCCGTCCGCGCC	2316 21082 2336 21022 2356 20968 2376	20908 GGTTGGCCTCCAGCCTGCAGCTGCGGCGAGAT	2453 AspCysAsnArgArgThrProLeuThrAsnArgValTrpGluAsp 2452
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	22169 TGCTGCGCGGGGGGCGTCAGGCCTTGGCTGCGACCGTCCTGGTTCCGCT 1955 rGlyHishlaPheLeualaLysProProAlaArgSerGlyLeuGluProAlaSerSerPr 122109 GACCCACGGACCACGCAAGCACACGCTCACGCTCGCGCGCGCGCTC 1975 oSerLysGlySerGluProArgProLeuValProProValSerGlyHisAlaThr11eAl	22001 CCAGAGAAGCTCTTGCAGCGGCCGTCTCGGGCCATCCCCTTGAGCCGGCTGAACTCCAG 1999 aLysAsnLeuAlaProHisHisAlaSerProAspProProAlaPro 21941 AACAACGCGGGGGTGCTCATCACCGTCACCCGGCCGGCCG	21764 CTGGCGGCTCCCGATGCCCCGGTACCCGTCCAGCATCTCCAGCCCGCCACCTGGTAA 2064 UThrHisAspLysGlyLeuProLysHisLeuGluGluLeuAspLysSerHisLeuGluGl 21704 GCGCCATAGTCCG	2122 rAlaProGlyvalLysGlyHisGlnArgValValThrLeuAlaGlnHisIleSerGluVa 21598 -GCACCAGCCGCTGTGGGATCCATCGCTGCGCGCGCTCTCGGGGGG 21598 -GCACCAGCCGCTGTGGGATCCATCGCTGCGCTCTCTCGGGGGG 2152 llleThrGlnAspTyrThrArgHisHisProGlnGlnLeuSerAlaProLeuProAlaPr 2153ATATCCCGAAGAAGTCCGCGTCGAACAGGTCGATGTCCCGCAAG 2162 oLeuTyrSerPheProGlyAlaSerCysProValLeuAspLeuArgArgProProSerAs ::::

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22-JAN-1998; 98US-00010809.
31-AUG-1998; 98US-00144085.
15-FEB-2001; 2001US-0271245P.
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       Polyangium cellulosum.
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                                                 ProGlyAsnThrSerGlnProProAlaPhePheSerLysLeuThrGluSerAsnSerAla 2281
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             recombinant DNA vector; tmbA gene cluster; tmbA; tmbB; tmbC;
polyketide synthase enzyme; PKS enzyme; hybrid PKS enzyme; tombamycin;
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           |||| || || : : : || || || ATGTTGTGAGCAAAGGAGTGATGATGATGATGA
                                                                          MetVallyaSerLysLysGlnGluIleAsnLysLysLeuAsnThrHisAsnArgAsnGlu
                                                                                                                                                                                            LeuGluAlaileileArgLysAlaLeuMetGlyLysTyrAspGlnTrpGluGlu-----
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TyrProleuLeuTyrArgAspGlyGluGlnThrGluProSerArgMetGlySerLysSer
                                                                                                                   ProGluTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetProAlaIleThrGly
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New recombinant DNA vector for preparing hybrid polyketide synthase enzymes comprises all or a portion of one or more genes in the tmbA gene cluster of Sorangium cellulosum.
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                                                                                                                              The invention describes a recombinant DNA vector that comprises all or portion of one or more genes in the tmbA gene cluster. Also disclosed in the host cell that comprises the tmbA genes of Sorangium cellulosum or fragments of those genes. The DNA vector is useful in preparing hybrid polyketide synthase (PKS) enzymes and the polyketides produced by such hybrid enzymes. The gene produces of the tmbA cluster can be used to synthesize the polyketide tombamycin. This sequence represents the Sorangium cellulosum tmbA gene cluster.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  28269 TTCTTCGCCACCGAGCTCGCCCCTCACGCCCCCACACGGGTACCTGCGGCCTCCCATA
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Retinoid X receptor (RXR) interacting protein (RIP) - useful to modulate or mediate RXR function, anti-RIP antibodies can be used to determine RIP
                                                                                                                                                                                                                                                                                                                                                                                                                                                 1024 CCTCCTGCTCTCGCCGCAGGCTCGCCACAAACGTCGCC------TCGGCTCCCT
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1603	HisHisProHisProIleSerProTyrGluHisLeuLeuArgGlyValSerGlyValA 1622
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1641	leproLeuAspAlaAlaAlaAyrTyrLeuProArgHisLeuAlaProAsnProThrT 1661
1661	yrProHisLeuTyrProProTyrLeuIleArgGlyTyrProAspThrAlaAlaLeuGluA 1681
1681	sAsnThrAl ccGcGccGG
1701	euArgGlyLeuSerProArgGluSerS
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polyketide synthase enzyme; PKS enzyme; hybrid PKS enzyme; tombamycin;
ds.
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8 8		ò	1008 GluSerAspAlaProGlnGlnProGlySerSerProA
Š 2	/36 IVEIONATAINEVALMENSELSELASPINICALUSELLISELLOSEL FELONELINGEN /3/	Db 1	19277 GCGCCGCCCACGCGCCCCGCGTCACCCACCAGG
3 8		λ	1028 ProProAlaAspLysGluAlaPheAlaAlaGluAlaG
) E	20246 GCTTGCGGAGATTCCAGGTGCACTGGACTCGCTTCAGGGCGTGAGAGCGCGGCGT 20187	Db 1	19217 ccaradcacectrecadcaccacaccac
8 8		δλ	1048 CysTrpThrSerGlyLeuProPheProValProProA:
: A	20186 CCAGCAGAGGCCCTCTTCCACCGGCATCGGCACCCATTGCGCCTCATGCGCG 20127	Db 1	19172CGTGGGACAACGACGCCACTCCCGACTG
t I		ò	1068 HisAlaProAspProSerAlaPheSerTyrAlaProP

LysglyLysAspAlaGluAlaAla 883 GCACAAGGTGCCTCACCCCGT--- 19611 ccadddcccccccddacacccdd 19398 GCCGGCGCTCGGGGTGCTCGCTGC 19338 ||| ||| ||| ccagarcercerce 19278 ------ccreerescrip 19173 |||:::||| |||| |GACTCTGTCCTAAAGCTGTCGTGT 19122 3GTGCGCCGCGAGCGCGTCGAGGA 20007 CGCCAAACGTACCGGCCGCCGACG 19947 CGGCCGTCTGGGCGGTGAGCACGC 19827 rcagegeegeegeegerge 19767 CGTCGCACGCACCGTCACCG 19707 |||| GACCCAGCTCCCTGTCCCGCCGG 19557 CICCICGGGIGAGCICITCCGCIG 19497 CGCCCCGCGCAGCGCAAGCTCCG 19437 CGCCCGCCTGTGTCCAGAAGCCCC 20067 ProProglnAsnLeuGlnPro 1007 GlnLysLeuProGlyAspProPro 1047 ArgGluvajilleLysAlaSerPro 1067 ValGluGluGlyGluGluGln--- 843 GlyLysAlaGluGluProValLys 863 AspSerSerAlaThrCysSerAla 923 Argieu--------- 938 AlaAlaAlaIleProProIleGln 977 ThrproproArg 787 LysGluGlyGlySerGlyArgAla 903 AspproArgAlaAsnAlaSerPro 957 AlaproProProValValProLys 824Gly

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comprising a PKS ORF (open reading frame) which encodes one or more pks domains is useful for producing polyketide synthases from which polyketides can be produced. The host cells are useful for constructing a polyketides can be produced. The host cells are useful for constructing a library, where each individual colony of the library represents a colony with the ability to produce a particular PKS synthase and ultimately a particular polyketides produced by these colonies can be used collectively in a panel to represent a library or may be assessed individually for activity. Colonies in the library are also induced to produce the relevant synthases and thus to produce the relevant colonies in the library are also induced to produce the relevant synthases and thus to produce the relevant colonies in the library are also induced to polyketides to obtain a library of candidate polyketides which can be screened for binding to desired targets such as receptors, signalling proteins, etc. The present sequence represents the DNA sequence of cosmid pKDS18-26 which encodes one or more domains of S. cellulosum PKS. Note:

The present sequence is said to encode the functional domains of S. cellulosum PKS which correspond to domains or domain subsets of the amino acid sequences of OKR1 (AMU10700) and OKF2 (AAU10701). (Updated on 11-SEP -2003 to standardise OS field)
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CGCGAACACCACCTCTCTCAACCCACGGTCGAGATACGCGTCCAGCGCCTCGCACACCTC 22360
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The present invention relates to the isolation of novel Sorangium cellulosum polyketide synthases (PKS), and the polynucleotide sequences encoding them. The polyketide synthases include catalytic domains such as ketosynthase domain, acyl transferase domain, dehydratase domain,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               LeuMetGluAlaLeuGluLysLysValGluArgIleGluAsnAsnProArgArgArgAla 324
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Novel purified, isolated DNA molecule from Sorangium cellulosum having polyketide open reading frame encoding modules with one or more domains such as ketosynthase, acyl transferase and acyl carrier protein domains.
                                                                                              298 AATGAGAAACAAATGCGGCAGCTCTCTGTGATTCCACCTATGATGTTTGATGCAGAACAA
                                                                                                                                                                                                                                                                                                                                                                            238 ATTGCTAGGAGTGAGCATGAGATTTCTGAAATTATTGATGGCTCTCTGAGCAGGAGAAT
                   61 CTCATGGAGGCATGGGAGAAAAAGTGGACAGAATAGAAAATAATCCTCGGAGGAAAAGCT
                                                                         LysGluSerLysValArgGluTyrTyrGluLysGlnPheProGluIleArgLysGlnArg
                                                                                                                                                                                        181 GAACAGCAAGAAAGATTTCAG---CGAGTTGGGCAGAGGGGAGCTGGTCTTTCAGCCACC
                                                                                                                                                                                                                           365 AlaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSerGluGlnGluAsn
                                                                                                                                                                                                                                                                                                      385 LeuGluLysGlnMetArgGlnLeuAlaValIleProProMetLeuTyrAspAlaAspGln
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ONA sequence of S. cellulosum polyketide synthase cosmid, pKOS28-26
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Polyketide synthase, PKS; catalytic domain; ketosynthase domain; acyl transferase domain; dehydratase domain; ketoreductase domain; acyl carrier protein domain; pKOS28-26; ds.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    445 MetGlnHisProLysAsnPheGlyLeulleAlaSerPheLeu 458
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AAS17367 standard; DNA; 33529
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(first entry)
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   The invention relates to polynucleotides encoding colon tumour proteins. The polynucleotides and encoded polypeptides are useful in pharmaceutical compositions, such as vaccines, for the diagnosis, prevention, and treatment of colon cancer. Polynucleotide sequences may be used as hybridisation probes or primers, and in the design and preparation of ribozyme molecules for inhibiting expression of tumour polypeptides and proteins in tumour cells. The compositions are useful for stimulating an immune response against cancer, particularly for the immunotherapy of colon cancer, and as markers for the progression of cancer. ABK4450-ABK4621. Tepresent coding sequences of human colon tumour proteins of the invention. Note: With the exception of SEQ ID No 1 and 2, the sequence data for this patent did not form part of the printed specification but was supplied by the European Patent Office
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      colon cancer polypeptides and polynucleotides, useful as vaccines, diagnosing, preventing, and treating colon cancer, and as markers for progression of cancer.
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                                                                                                                                                                                                                                                                                                      colon tumour; vaccine; colon cancer; immunogenic; immunotherapy;
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                                   ArgileGluasnabnProargargargalalysGluSerLysValarg
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                    GinLeuAlaValileProProMetLeuTyrAspAlaAspGinGlnArgIleLysPheile
                                                              AsnMetAsnGlyLeuMetAlaAspProMetLysValTyrLysAspArgGlnValMetAsn
                                                                                    AACATGAATGGGCTTTATGGAGGACCCTATGAAGTGTATAAAGATAGGCAGTTTATGAAT
                                                                                                                     PheGlyLeuIleAlaSerPheLeuGluArgLysThrValAlaGluCysValLeuTyrTyr
                                                                                                                                                                                                           GAGATTTCTGAAATTATTGATGGGCTCTCTGAGCAGGAGAATAATGAGAAACAAATGCGG
                                                                                                           MetTrpSerGluGluLysGluThrPheArgGluLysPheMetGlnHisProLysAsn
                                                                                                                                                                                               TyrLeuThrLysLysAsnGluAsnTyrLysSerLeuValArgArgSerTyr---ArgArg
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                                         caectrororoganticoacctangangitudanecaeaacaaagaceagtcaagitcarr
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Human; ss; gene; prey; adipocyte; SID; selected interacting domain; anorectic; antidiabetic; protein-protein interaction; diabetes; yeast 2-hybrid assay; metabolic disorder; obesity.
                                                                                                                                                                                                                                                                                                                                                                                                                                                              Human adipocyte Selected Interacting domain, SID, cDNA #610
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by selecting a bait protein from a known adipocyte marker and then performing a yeast 2-hybrid selection to isolate prey proteins are designated by members of an adipocyte confar library. The proteins are designated by members of an adipocyte confar library. The proteins are designated by members of an adipocyte colls. Also included are a polymerical interacting domains) proteins. Also included are a polymerical encoding a polypeptide in the adipocyte cells, a recombinant host cell expressing at least one of the interacting collypeptides of the comprising a modulating compound in adipocyte cells, a SID (RTM) polypeptide comprising any of the 738 amino acid sequences given in the specification (including its fragment or variant), a SID (RTM) polymerical comprising any of the 738 amino acid signen in the specification (including its fragment or variant), comprising the SID (RTM) polymerical and its fragment or variant), comprising the vector, a protein chip comprising the polymerides and a recombinant host cell crecord comprising all or part of the data, listed in the specification. The complex, polymerical disorders such as one diabetes.

The complex, polymerides, polymerical disorders such as obsetty or diabetes. are identified and then The polymucleotides are useful as probes or primers. The complex is particularly useful for identifying selected interacting domains (SID (RTW)) for screening drugs that medulate the protein interaction, thus exhibiting the therapeutic effect. The present sequence encodes a SID given in the specification. (prey) protein of the invention

Sequence 673 BP; 245 A; 134 C; 153 G; 141 T; 0 U; 0 Other;

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AAGAAACAACAACAGCTTGAAGAAGAGGCAGCTAAAACCTCCTGAGCCCTGAGAAGCCCGTG 325
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LysGlnLysPheCysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGlu 314

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The invention relates to a complex between two interacting proteins

Claim 7; Page 308-309; 382pp; English.

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complementary to the specified genes. The device of the invention may be used to determine gene expression profiles for measurement of monocyte or macrophage activation or inflammation in blood or other tissues, as well as for the diagnosis, prognosis and monitoring therapy in cases of rheumatoid arthritis, chronic inflammatory disease, chronic bacterialinduced inflammation, arteriosclerosis, tumours, organ or tissue transplants and sepsis. The device provides information for molecular classification and staging of disease, creation of a patient-specific prognostic profile and suggestions about molecular pathogenesis or therapeutic effects, thus facilitating the development of new therapeutic strategies and pharmacological concepts. The current sequence is that of
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BP; 91 A; 229 C; 180 G; 204 T; 0 U; 14 Other;
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The invention relates to novel genes (AAI93926-AAI97963) expressed in human neuroblastoma. The nucleic acids are applicable as a probe or primer in diagnosing the prognosis of human neuroblastoma, malignancy and susceptibility indicators or tumour markers for anti-cancer agents. The gene information for diagnosing prognosis is related to factors similar to that for N-myc and TrkA genes
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    GluTyrTyrGluLysGlnPheProGluIleArgLysGlnArgGluLeuGlnGluArgMet 350
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                                                                                                                                                                                                                                                                                                                                                                                 Nucleic acids originating in gene expressed in human neuroblastoma, useful as probe or primer in diagnosing prognosis of human neuroblastoma, malignancy and susceptibility indicator or tumor marker for anti-cancer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ArgiysGlnTrpLysGlnLysPheCysGlnArgTyrAspGlnLeuMetGluAlaLeuGlu
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   neuroblastoma; malignancy; cancer; tumour marker; N-myc;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Sequence 710 BP; 145 A; 173 C; 107 G; 282 T; 0 U; 3 Other;
                                                                                                       SEQ ID NO
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Matches:
Conservative:
Mismatches:
Indels:
                                                                                                       expressed polynucleotide
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                                                                                                                                                                                                                                                                                                                                                                                                                                                   Claim 1; Page 2623; 2979pp; Japanese.
                           BP.
                                                                                                                                                                                                                                                                                       CHIBA PREFECTURE.
HISAMITSU PHARM CO LTD.
                           AAI97539 standard; cDNA; 710
                                                                                                                                                                                                                                       02-MAR-2001; 2001WO-JP001629
                                                                                                                                                                                                                                                               07-MAR-2000; 2000JP-00159195
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845.00
82.28%
68.50%
6.39%
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                                                                                                       Human neuroblastoma
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Best Local Similarity:
                                                                                                                                                                                   WO200166719-A1
                                                                                                                                                                                                                                                                                                                                 Nakagawara A;
                                                                                                                                                           Homo sapiens.
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Pred. No.:
                                                                             13-NOV-2001
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                                                   AAI97539,
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DB:
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RESULT 17
AAI97539/c
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GluValSerGluIleIleAspGlyLeuSerGluGlnGluAsnLeuGluLysGlnMetArg 390

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370 476

351 GlnSerArgValGlyGlnArgGlySerGlyLeuSerMetSerAlaAlaArgSerGluHis

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The inversion trainers to a complex between to interacting jucturing in the inversion trainers to a complex between the proteins are identified by selecting a bait protein from a known adipocyte marker and then be reforming a yeast 2-hybrid selection to isolate prey proteins encoded by members of an adipocyte cDNA library. The proteins are designated SID (RTM) (selected interacting domains) proteins. Also included are a polymeptide encoding a polypeptide in the adipocyte cells, a compliant host cell expressing at least one of the interacting complex, selecting a modulating compound in adipocyte cells, a SID (RTM) polypeptide comprising any of the 738 amino acid sequences given in the specification (including its fragment or variant), a SID (RTM) polymucleotide comprising any of the 738 nucleotide sequences given in the specification (including its fragment or variant), a vector comprising the SID (RTM) polymucleotide, a recombinant host cell comprising the vector, a protein chip comprising the polypeptides and a recombinant host cell comprising all or part of the data, listed in the specification. The complex, polypeptides, polymucleotides and compounds are useful for the complex, polypeptides, polymucleotides and compounds are useful for the complex properties and compounds are useful for the part of the data, listed in the specification.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     New complex between two interacting proteins in adipocyte cells, useful for identifying selected interacting domains that modulate protein interactions, or for preventing or treating metabolic disorders such as obesity or diabetes.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    The polynucleotides are useful as probes or primers. The complex is particularly useful for identifying selected interacting domains (SID (RTM)) for screening drugs that medulate the protein interaction, thus exhibiting the therapeutic effect. The present sequence encodes a SID
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 invention relates to a complex between two interacting proteins in
                                           Human, ss; gene, prey, adipocyte, SID, selected interacting domain, anorectic, antidiabetic, protein-protein interaction, diabetes; yeast 2-hybrid assay, metabolic disorder, obesity.
Human adipocyte Selected Interacting domain, SID, cDNA #611.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Claim 7; Page 309; 382pp; English.
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                                                                                                                                                                                                                                                                                                  14-MAR-2002; 2002WO-EP003768
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P-PSDB; ABU70980.
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                                                                                                                                                     Homo sapiens
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Sequence 555 BP; 151 A; 162 C; 167 G; 75 T; 0 U; 0 Other; 0000 Length:
Matches:
Conservative:
Mismatches:
Indels: Gaps: 6.33e-22 960.00 100.00% 100.00% 7.26% Percent Similarity: Best Local Similarity: Alignment Scores: Query Match: DB: Score:

AspArgSerLeuThrGlyLysLeuGluProValSerProProSerProProHisThrAsp US-09-522-753-5 (1-2517) x ACA57524 (1-555) 138

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177 cercadecrecadecrecedecedecrerereadecadecrearecadaacarecadad 121 61 ProGluLeuGluLeuValProProArgLeuSerLysGluGluLeuIleGlnAsnMetAsp GACCGTAGCCTGACGGGCAAGCTGGAACCGGTGTCTCCCCCCCAGCCCCCCGCACACTGAC ~ 158 62

Device for diagnosis, prognosis and monitoring therapy of e.g. rheumatoid arthritis, comprises immobilized gene sequences from monocyte-macrophage The invention relates to a novel device for diagnosis, prognosis and monitoring of therapy having, on its surface, sequences of some or all of specified monocyte-macrophage genes and also of other genes or RNA 217 237 277 297 181 241 301 257 481 PheCysGlnArgTyrAspGlnLeuMetGluAlaLeuGluLysLysValGluArgIleGlu 317 TTCTGCCAGCGCTATGACCAGCTCATGGAGCCTTGGAAAAAAAGGTGGAGCGCATCGAA 541 ArgValAspArgGluIleThrMetValGluGlnGlnIleSerLysLeuLysLysGln 197 actreaaccrecacarcedarreredaadecerdedececeaderecacereceererae 361 AACCAGCCTCCGACACCCGGCAGTATCATGAGAACATCAAAATAAACCAGGCGATGCGG 421 CAACAGCTGGAGGAGGAGGAGCTGCCAAGCCGCCCGAGCCTGAGAAGCCCGTGTCACCCGC ProlleGluSerLysHisArgSerLeuValGlnIleIleTyrAspGluAsnArgLys cccarcaagregaagcaccagccregregagarcarcracaacaagaagcagaagaag AAGAAGCTAATCTTGTACTTCAAGAGGAGGAATCACGCTCGGAAACAATGGAAGCAGAAG GinglinLeuGluGluAlaAlaLysProProGluProGluLysProValSerProPro AlaGluAlaAlaHisArgIleLeuGluGlyLeuGlyProGlnValGluLeuProLeuTyr AsnGlnProSerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArg LysLysLeulleLeuTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLys monocyte, macrophage; gene expression profile; rheumatoid arthritis; chronic inflammatory disease; bacteria-induced inflammation; arteriosclerosis; tumour; organ; tissue transplant; sepsis; molecular classification; human; BSK; ss. SEQ ID NO 171; 180pp; German. BP 02-OCT-2002; 2002EP-00090348 09-NOV-2001; 2001DE-01055600 ADE76358 standard; DNA; 718 Ë AsnAsnProArg 321 AACAACCCGCGC 553 Haeupl (OLIG-) OLIGENE GMBH WPI; 2003-443090/42. (first Human BSK-23 DNA Stuhlmueller B, Homo sapiens EP1310567-A2 29-JAN-2004 14-MAY-2003 Claim 29; 482 178 198 182 218 242 238 302 362 278 422 298 318 542 258 ADE76358; RESULT 16 ADE7638/I ID ADE7/ AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 AC ADE7 g 원 8 g Š g à g ð ద ò g à g ò ઠ

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                                                                                                                               Human, colon cancer; tumour, diagnosis, gene expression product; probe; detection; cancerous state; metastasis; identification; breast cancer;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         mammalian
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 metastatic or metastatic cancerous states, stages of cancer, or responsiveness of cancer to therapy). This is particularly for breast cancer, oestrogen receptor-positive breast cancer, oestrogen receptor-
                                                                                                                                                                                                                                                                                                                                                                                                                             A, Garcia PD, Sudduth-Klinger J;
Kennedy GC, Pot D, Kassam A;
R, Dickson M, Drmanac S, Labat
Jones LW, Stache-Crain B;
                                                                                                       colon cancer cell line polynucleotide sequence SEQ ID NO:2661
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        cancerous states of
                                                                                                                                             detection; cancerous state; metastasis; identification; bre oestrogen receptor-positive breast cancer; therapy; oestrogen receptor-negative breast cancer; lung cancer; ss
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Matches:
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Siese K, Randazzo F, K
Manac R, Crkvenjakov R,
Kita D, Garcia V, Jo
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                          BP
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98US-0085337P.
98US-0085696P.
98US-0105234P.
98US-0105877P.
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1076.00
82.62%
70.82%
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                        AAA02670 standard; cDNA; 872
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Drmanac R,
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Leshkowitz D, Kita D,
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HYSEQ INC.
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Reinhard C,
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15-MAY-1998;
15-MAY-1998;
21-OCT-1998;
27-OCT-1998;
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Pred. No.:
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                                                    AAA02670;
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                                                                                                       Human
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Conservative: Mismatches: Indels:

Percent Similarity: Best Local Similarity:

Query Match: DB:

Gaps:

US-09-522-753-5 (1-2517) x AAA02670 (1-872)

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                                                            SerAspThrArgGlnTyrHisGluAsnIleLysIleAsnGlnAlaMetArgLysLeu
                                                                                281 IleLeuTyrPheLysArgArgAsnHisAlaArgLysGlnTrpLysGlnLysPheCysGln
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                                                                                                                                                                                                          ArgargargalalysGluSerLysValargGluTyrTyr-GluLysGlnPheProGluIl
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                                                                                                                                                                                                                                                                                                                                                   310 rcgadadcadadadadcadcadadadatrrcag---cgagrrcggcagaggggggcrcg
                                                                                                                                                                                                                                                                                                                                                                                 360 yLeuSerMetSerAlaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSe
                                                                                                                                                                                                                                                                                                                                                                                                   rGluGlnGluAsnLeuGluLysGlnMetArgGlnLeuAlaVallleProProMetLeuTy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                427 TGAGCAGGAGAATAATGAGAACAAATGCGGCAGCTCTC-GTGATTCCACCTATGATGTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  tLysValTyrLysAspArgGlnValMetAsnMetTrpSerGluGluLysGluThrPh
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                546 GAAAGTGTATAAAGATAGGCAGTTTATGAATGTTTGGACTGACCATGAAAAGGAGATCTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             440 eArgGluLysPheMetGlnHisProLysAsnPheGlyLeuIleAlaSerPheLeuGluAr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               SSerleuValArgArgSerTyr---ArgArgArgGlyLysSerGlnGlnGlnGlnGlnGl
|:::|||||||||||::|||
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       726 AGCCCTCGTCAGAAGGAATTATGGGAAACGCAGAGGCAGAAACCAGCAA-----
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      519 ulysAspGluLysGluLysGluLysGluLysGluAlaGluLysGluGluLysProGluValGl
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        499 nGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnGlnProMetProArgSerSerGlnGluGl
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අ <u>ධ</u>	CAC	Qy 2221	
<u></u> 음		Db 10090	GGGCGCAACATAATCCG
ò	CysProLeuGlyGlyThrLeuAspGlyValTyrProThrLeuMetGluProValLeuLeu	1	
q		Qy 2249	G1G1
<i>&</i> 4	1939 ProLysGluAlaProArgValAlaArgProGluArgProArgAlaAspThrGlyHisAla 1958 1939 ProLysGluAlaProArgValAlaArgProGluArgProArgAlaAspThrCArgCargCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargCargTcArgCargTcArgCargTcArgCargTcArgCargTcArgCargCargCargCargCargTcArgCargCargCargCargCargCargCargCargCargCa	1	GCGGGTGGTGAAG
g &	GGTCGC		
S 8	CGGTTGCCGCCACGAGTTCCGGATCGGGAGGCGGCACCGGTACACGATCCTCA	Db 10240	GGAACGCCCAGCGGTGC
ò	Pro	Db 10300	
ପ୍ପ			GlulleAsniyşiysie
ò		Db 10360	 GGTTATGGAAAGAATCC
අධ		Qy 2309	ProGlyThrGluIleP
δ	ProAlaProProAlaSerAlaSerAsp	Db 10408	
q	CCCAGTGTGATTAATATTGATTTGGATCAGGAGCGCATCTCAGCGGCAGCAGCAGTT	Qy 2329	SerGlnAlaValGlnG
ò	ProHisArgGluLysThrGlnSerLysProPheSerIleGlnGluLeuGluLeuArgSer	Db 10429	10429 TCGCATGCCGCCG7
qq		Qy 2349	AlaLeuMetGlyLysT)
ò	GlyTyrHisGlySerSerTyrSerProGluGlyValGluProValSerProValSer :::	Db 1048	 10486 GCTGCTATGGAGTTGC2
qq		Qy 2363	
ò	SerproSerLeuThrHisAspLysGlyLeuProLysHisLeuGluGluLeuAspLysSer :::	Db 10546	CGTAAGACGCCCAACC
q	AGTCCACAGAGATTCATACGAAGGCATTACCTTTGGGGAACTAACCGATTCG	Qy 2382	2 ProlleThrAlaAlaA
ò	<pre>HisLeuGluGlyGluLeuArgProLysGlnProGlyProValLysLeuGlyGlyGluAla :::</pre>	0901 qa	10606 CCTGGTGCCTTTGCCAA
qq		Qy 2400	0 GlyGlyLysAlaLysV
ò	AlaHisLeuproHisLeuArgPro	Db 10666	GGTGGGTGTCATTGG
qq		Oy 2420	0 GlyLeuAlaSerGlyA
ò	SerSerPro	10702	2 GGAATTGCAGGCGGA-
qu	CGGATCGCTGGAAGCAGAACGTCGTATGCAACAGAAAGGCCGAGGAAGCGAAG	Qy 2440	0 AsnArgArgThrProL
ò	GlyHisGlnArgValValThrLeuAlaGlnHisIleSerGluValIle	Db 10731	1
qq		Qy 2460	0 ThrProPheProTyrA
ò	2144 ThrGlnAspTyrThrArgHisHisProGlnGlnLeuSerAlaProLeuProAlaPro 2162	Db 10732	
qq	9931 CAGCAGCAGCAGCAGCACACACCAGCAACATCATCCGCAGATGCCGGGCACTGGC 9990	Qy 2480	
ò	2163 LeuTyrSerPheProGlyAlaSerCysProValLeuAspLeuArgArgProProSerAsp 2182	Db 10741	::: 1
qq	9991 TCTGGCTCGGCTCCAGGTGCT10014	Qy 2495	5 HigHis
ò		Db 10792	::: 2 CACAATTCCAGTTCAT
qq	10015GGTCAGGCGGTGGATCGGCCGCCCCGGGTAGCGGTGGT 10053	Oy 250	0 GlugiuProLysproL
ò	AsnLysThrSerValLeuGlyGlyGlyGluAsp		 10852 AGTGAGTCCAAGCCGC
QQ	10054 GGTGGCGCCGGCAGAGCCAGCACACCT	_	

10090	GlylleGluProValSerProProGluGlyMetThrGluProGlyHisSerArgSerAla 2240 ::
2241	ValTyrProLeuLeuTyrArgAsp 2248
10120	
2249	257
10180	102
2258	8
10240	
2269	œ
10300	
2289	GluileAsnLysLysLeuAsnThrHisAsnArgAsnGluProGluTyrAsnIleSerGln 2308
10360	GGTTATGGAAAGAATCGGACTGTGGGAACAGAGGATGAAGTA 10407
2309	ProGlyThrGluIlePheAsnMetProAlaIleThrGlyThrGlyLeuMetThrTyrArg 2328
10408	àccgccgaràrdgrgccarar 10428
2329	SerGlnAlaValGlnGluHisAlaSerThrAsnMetGlyLeuGluAlallelleArgLys 2348
10429	
2349	AlaLeuMetGlyLysTyrAspGlnTrpGluGluSerProPro2362
10486	
2363	LeuSerAlaAsnAlaPheAsnProLeuAsnAlaSerAlaSerLeuProAlaAlaMet 2381
10546	CGTAAGACGCCCAACCAATACGAGGTGGTAGACGCCCAGTGGCGACCAGGCGGTGCCA 10605
2382	ProlleThrAlaAlaAspGlyArgSerAspHisThrLeuThrSerProGlyGly 2399
2400	GlyGlyLysAlaLysValSerGlyArgProSerSerArgLysAlaLysSerProAlaPro 2419
10666	:::
2420	GlyLeuAlaSerGlyAspArgProProSerValSerSerValHisSerGluGlyAspCys 2439
10702	::: GGAATTGCAGGCGGACCAGGTGGTGTTCTCA
2440	
10731	10731
2460	ThrProPheProTyrAsnProLeuIleMetArgLeuGlnAlaGlyValMet.
10732	10740
2480	ProprogroGlyLeuProAlaGlySerGlyProLeuAlaGlyPro 2494
10741	GGGGAGGGGGGGGGGGGGGGGGGGGGGGGGGG
2495	HisHisAlaTrpAsp 2499
10792	
0 6	GluGluProLysProLeuLeuCysSerGlnTyrGluThrLeuSerAspSerGlu 2517
10852	AGTGAGTCCAAGCCGCTGCTGTTCCAAGTACGATGCACTCAGCGACGACGAAGAATTTCAG

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SerArgleuAspArgGlyArgGl 1260 CAAACATTGGACTTGAGCATCAA 7157 LysLysGlyHisValLeuSerTy 1280 AspGlyArgSerSerSer---- 1298 |||::: |||::: --ProAlaProProAlaAspLys 1032 luHisAlaLysAlaProValG| 1146 ArgGlyThrAlaLeuGlySerVa 1203 ||| ::: :::|| CCGGAGACCATAATCTACAATGT 6986 .----CCGTCGCAGCACTCGGT 7037 ProAlaAspValLeuTyrLysGl 1241 CATCCGCAGCATTCGTCGCATGG 7097 ------ArgThrTyrAspMe 1313 sluArgGlnIleGlyAlaIleSe 1126 CCACCAGCACTGGATGTGCGGG 6746 ::: || rcggtggtcaattcgcacatggg.6806 ystysteudlaProPheSergl 1166 SCAAGTTCCGCCGGAGGAATCGG 6866 SinAlaGlyProProGluSerLe 1184 :::||| rctagrggatcatcgaagagcgg 6926 ArgvalproSerAspSerAlall 1223 -----CGTTCGCTGAAG 6378 nProGluSerAspAlaProGln 1013 pProProCysTrpThrSerGly 1052 AGGAGCTTACCATTGTAAGGGA 6557 laSerProHis----- 1068 AGCAGCAGCAGCAGCAGGC 6608 roProGlyHisProLeuProLe 1086 CGCATGGC-------6654 roProThrileSerAsnProPr 1106 CAGCTGCAGTGGCACCACCGCC 6704 gGluAspAlaAlaProThrLys 993

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                                                                                               CAGCGCGAGCTGCAGGAGCGCATGCAGAGCAGGGTGGGCCAGCGGGGGCAGTGGGCTGTCC 481
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             The invention relates to an isolated nucleic acid detection reagent capable of detecting 1000 or more genes from Drosophila. The invention useful in developmental biology and in elucidating cell signalling and cell-cell interactions in higher eukaryotes for the development of insecticides, therapeutics and pharmaceutical drugs. The invention discloses genomic DNA sequences (ABL16176-ABL30511), expressed DNA ABG72072). The sequence data for this patent did not form part of the printed specification, but was obtained in electronic format directly
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   detection reagent for detecting 1000 or more for elucidating cell signaling and cell-cell
                         MetSerAlaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSerGluGln
                                          GluasnLeuGluLysGlnMetArgGlnLeuAlaValIleProProMetLeuTyrAspAla
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Sequence 10910 BP; 2875 A; 3179 C; 3157 G; 1699 T; 0 U; 0 Other;
                                                                                                                                                                                                                                                                                                    Drosophila melanogaster expressed polynucleotide SEQ ID NO 3875.
                                                                                                                                                                                                                                                                                                                             Drosophila, developmental biology; cell signalling; insecticide
                                                                                                                                      AspGlnGlnArgileLysPheileAsnMetAsnGlyLeuMetAlaAsp 418
                                                                                                                                                                 649
                                                                                                                                                    GACCAGCAGCATCAAGTTCATCAACATGAACGGGCTTATGGCCGAC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Claim 1; SEQ ID NO 3875; 21pp + Sequence Listing; English
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ftp.wipo.int/pub/published_pct_sequences
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11-JUL-2000; 2000US-00614150
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   New isolated nucleic acid ogenes from Drosophila and interactions.
                                                                                                                                                                                                                                                                            (first entry)
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                                                                                                                                                                                                                                                                                                                                                 pharmaceutical; gene; ss.
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P-PSDB; ABBS9028.
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      GluLeuGluLeuValProProArgLeuSerLysGluGluLeuIleGlnAsnMetAspArg 178
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     199 GlnLeuGluGluAla-----AlaLysProProGluProGluLysProValSer---
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                                                                                                                                                                                                                                                                                                                                                                                       ProProHisSerLeuSerTyrProValGinIleAlaArgThrHisThrAspValGlyLeu
                                         2092 CCACACACAC------CCGCAGCAG------CAACACCAGCAGCAGCAG
                                                                                        40 LeuGluTyrGlnHis-----HisSerArgAspTyrAlaSerHisLeuSerProGlySer
                                                                                                                                                                                                                                                                     -----LeuLeuSerGluPheGlnProGlyAsn
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Length:
Matches:
Conservative:
Mismatches:
Indels:

1.07e-24 1083.00 33.03% 22.06% 8.20%

Local Similarity:

Best

Query Match:

Percent Similarity:

Alignment Scores:

US-09-522-753-5 (1-2517) x ABL03131 (1-10910)

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633 AATGAGCCTGAATACAATATCAGCCAGCCTGGGACGGAGATCTTCAATATGCCCGCCATC 574
                                                                                                                             ATGGGGCTGGAGGCCATAATTAGAAAGGCACTCATGGGTAAATATGACCAGTGGGAAGAG 470
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                                                 -----AGAAGCCAGGCGGTGCAGGAACATGCCAGCACAAC
                                                                                                                                                             SerProProLeuSerAlaAsnAlaPheAsnProLeuAsnAlaSerAlaSerLeuProAla
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          2460 ThrProPheProTyrAsnProLeuIleMetArgLeuGlnAlaGlyValMetAlaSerPro
                                                                                                                                                                              469 TCCCCGCCGCTCAGCGCCAATGCTTTTAACCCTCTGAATGCCCAGTGCCAGCCTGCCCGCT
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                                ThrGlyThrGlyLeuMetThrTyrArgSerGlnAlaValGlnGluHisAlaSerThrAsn
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                                                                                                                                                                                                                                                                                                                         349 GGCGGGAAGGCCAAGGTCTCTGGCAGACCCAGCAGCCGAAAAGCCAAGCCCCAGCC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     Human adipocyte Selected Interacting domain, SID, cDNA #488.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       2500 GluGluProLysProLeuLeuCysSerGlnTyrGluThrLeuSer 2514
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      112 ACTCAAAGTAAACCCTTTTCCATCCAGGAACTGGAACTCCGTTCT 68
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                                                              573 ACCGGAAC----
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P-PSDB; ABU70857.
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The invention relates to a complex between two interacting proteins in adjocyte cells, given in the specification. The proteins are identified by selecting a bait protein from a known adjocyte marker and then performing a yeast 2-hybrid selection to isolate prey proteins encoded by members of an adjocyte cDNA library. The proteins are designated SID (RTW) (selected interacting domains) proteins. Also included are a polymedies of the complex, selecting a modulating compound in adjocyte cells, a recombinant host cell expressing at least one of the interacting cells, a SID (RTW) polymeptide comprising any of the 738 amino acid sequences given in the specification (including its fragment or variant), a SID (RTW) polymelectide comprising any of the 738 mucleotide sequences given in the specification (including its fragment or variant), a vector comprising the SID (RTW) polymucleotide, a recombinant host cell comprising the vector, a protein chip comprising the polypeptides and compounds are useful for the complex, polypeptides, polymelectides and compounds are useful for many control of the cata, listed in the specification of the cata, listed in the specification. New complex between two interacting proteins in adipocyte cells, useful for identifying selected interacting domains that modulate protein interactions, or for preventing or treating metabolic disorders such as The polymucleotides are useful as probes or primers. The complex is particularly useful for identifying selected interacting domains (SID (RTM)) for screening drugs that medulate the protein interaction, thus exhibiting the therapeutic effect. The present sequence encodes a SID 7; Page 271; 382pp; English. (prey) protein of the invention or diabetes obesity Claim

Sequence 650 BP; 170 A; 186 C; 205 G; 89 T; 0 U; 0 Other;

Alignment Sco	Scores:				
Pred. No.: Score: Percent Simi Best Local S Query Match:	larit imile	1.48e-26 1113.00 100.00% 100.00% 8.42%	Length: Matches: Conservative: Mismatches:	650	
-09-522	-753-5 (1-2517)	, × ACA57401 (1	-650)	•	
, Ya	203 GluAlaAla	LysProProGluProG	3luLysproValSerP	GlualaalaLysProProGluProGluLysProValSerProProProlleGluSerLys 222	2
Q	2 GAGGCTGCC	AAGCCGCCCGAGCCTG	BAGAAGCCCGTGTCAC		
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qo	62 CACCGCAGC	CTGGTGCAGATCATCT	racgacgagaaccgga	CACCGCAGCTICITITITITITITITITITITITITITITITITITI	121
, va	243 ArgileLeu	ArgileLeuGluGlyLeuGlyProGlnValGluLeuProLeuT	31nValGluLeuProL	rAsnGlnProSerAsp	262
Db 1	122 CGGATTCTG	GAAGGCCTGGGGCCCC	CAGGTGGAGCTGCCGC		181
,, ,,	263 ThrArgGlnTyrHi	TyrHisGluAsnIleLys	slleAsnGlnAl	aMetArgLysLysLeulleLeu 28	282
Db 1	182 ACCCGGCAG	HILLIIIIII	AAAATAAACCAGGCGA	ACCCGGCAGTATCATGAGAACATCAAAATAAACCAGGCGATGCGGAAGAAGCTAATCTTG 241	=
"	283 TyrPheLys	sArgArgAsnHisAlaA	aArgbysGlnTrpLysG	sGlnLysPheCysGlnArgTyr 30	302
° qo	242 TACTTCAAG	AGGAGGAATCACGCTC	CGGAAACAATGGAAGC		301
č	303 AspGlnLeu	AspGlnLeuMetGluAlaLeuGluLy	ystysvalGluArgi	sLysValGluArglleGluAsnAsnProArgArg 32	322
0	302 GACCAGCTC	ATGGAGGCCTTGGAAA	AAAAGGTGGAGCGCA	GACCAGCTCATGGAGGCCTTGGAAAAAAGGTGGAGGGCATCGAAAACAACCGCGCGGG 361	11
à	323 ArgAlaLys	ArgalaLysGluSerLysValArgGluT	YFTY	rGlubysGlnPheProGlulleArgbys 342	C)
qq	362 CGGCCAAG	CGGGCCAAGGAGCAAGGTGCCCGAGTACTACGAAAAGCAGTTCC	SAGTACTACGAAAAGC	AGTICCCTGAGATCCGCAAG 421	=

343 GlnArgGluLeuGlnGluArgMetGlnSerArgValGlyGlnArgGlySerGlyLeuSer 362

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cancer) or other call proliferative disorders (e.g. arteriosclerosis, atherosclerosis, cirrhosis, hepatitis, polycythemia vera, primary thrombocytopenia, autoimmunia, polycythemia vera, primary thrombocytopenia, autoimmunia, polycythemia vera, primary thrombocytopenia, autoimmunia, finlanmatory disorders (e.g. Addison's disease, thyroiditis, multiple sclerosis, osteoarthritis, rheumatoid thanhimoto's thyroiditis, multiple sclerosis, osteoarthritis, rheumatoid arthritis, scleroderms, systemic lupus erythematosus), infections (e.g. viral, bacterial, fungal or parasitic infection), developmental disorders (e.g. thrombosis, hypopituitarism, hypogonadism, gigantism, golter) metabolic disorders (e.g. hypercholesterolaemia, hypogivalemia, diabetes, chyperlipidaemia, obesity), neurological disorders (e.g. ischaemic creebrovascular disease, stroke, Alzheimer's disease, pick's disease, Huntington's disease, parkinson's disease, Pick's disease, anxiety, schizophrenia), gastrointestinal disorders (e.g. ulcers), transport disorders (e.g. akinesia or multidrug resistance), or connective tissue disorders (e.g. Paget's disease or rickets). This cof the invention.
cervix cancer, bone cancer, liver cancer, lung
           cancer,
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Sequence 2336 BP; 367 A; 634 C; 894 G; 441 T; 0 U; 0 Other;

ores: 2 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	3-5 (1-2517) x ADE31306 (1	ProValLeuLeuLeuproLysGluAlaProArgValAlaArgProGluArgProArgAlaAsp 1954	5 ThrGlyHisAlaPheLeuAlaLysProProAlaArgSerGlyLeuGluProAlaSerSer 1974	s ccarcrcacecreecricece	ProserLysgly-SerGluProArgProLeuValProProValSerGlyHisAl 199	34 CCTICITGCCTAATCTCTCTCTGTGTCTCCTCTCTGTCTGTCTGTC	e~ ~	Arcrereacerecerregeereee	04	54 IGTCTTIGGGAGCCCAAACCCIACCCIAGCTIGGGTÍCCCCÍTGÁCCCCCCGGGGGTCC 2005	15 roAlaSerAlaSerAspProHisArgGluLysThrGlnSerLysProPheS 2032	1945 CAGCCAGCTGGGAGGGCAGCCCTGCCCTCGGAACCCTGGGCCCGGTGCCTGA 1945	32 erileGlnGluLeuGluLeuArgSerLeuGlyTyrHisGlySerSerTyrSerProGluG 2052	CTCTGCACCCCCGCCT		90 degresadecedereadecereridaderecadarereacedadadadadecede 1831	72 ysHisLeuGluGluLeuAspLysSerHisLeuGluGluLeuArgProLysGlnProG 2092	30 AGCACCTGGAAGAGCTCGACAAGAGCCACCTGGAGGGGGGGG	92 lyproValLysLeuGlyGlyGluAlaAlaHisLeuProHisLeuArgProLeuProGluS 2112	70 gececengaagerngsegssasassasseseseseseseseseseseseseses	12 erGlnProSerSerSerProLeuLeuGlnThrAlaProGlyValLy8GlyHisGlnArgV 2132
nment S No.: e: ent Sim Local y Match	-09-522-75	1935	1955	223		218	199	2124	2004	2064	2015	2004	2032	1944	2052	1890	2072	183	2092	1770	2112
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Db 171	0 GCCAGCCCTCGTCCAGCCGCTGCTCCAGACCGCCCCAGGGGTCAAAGGTCACCAGGGG 1651
213	2 alvalThrLeuAlaGlnHisIleSer 2140
165	0
214	.0 2140
Db 159	O TIGCACCIGGGGGCACCAGGCCIGGAGAGGGGAIGGGGAACCCCACAGCCCTICIGICCT 1531
214	.0 2140
Db 153	O GCCGGGGTGGCTGGGGGATCCAGGGCATGGCGCTGGGGGGATCCAGGGCGTGGGTGAGGG 1471
214	.0 2140
Db 147	10 TGAGATCCCAAAGCCCCGAGCACCGGCACCATCACCGCCCCCTAATCCATGGGAGGAGCC 1411
214	10 2140
Db 141	.0 TGTGATGCGAGCCGATGGCATCTTCACGGGCAATGAGGCCTTCCTGGTGGCCCAGGTTTC 1351
Qy 214	10 2140
Db 139	50 TCAGTGTCATGGGCTGGTCTCATCAGCCATCTGCCAACTACCAGCTTGGGACCGCTGACC 1291
Oy 214	40 2140
Db 129	90 ACAGCCCCACTCCCATGCACACTGGGACACGGGGCCCCAGAGGTGGCGGCGGCAGGTCCAC 1231
21,	40 2140
Db 12;	30 AGTCACCCAGGAAGCTGGCCCCACCAGGATTCTGCCCCGAGGCTCCGTCTAGCCCCTCCC 1171
0y 21,	40 2140
Ob 11.	70 CACCCCCAGAAGGTTCTGTCAGGAGAGTGCTGCCTGACTCTGGGCCCCCCCC
Dy 21,	15
0b 111	o caggadercarcacacacacacacacacacacacacacacacacac
2y 216	0 ProAlaProLeuTyrSerPheProGlyAlaSerCysProValLeuAspLeuArgArgPro 21
Db 105	o cacccccctrarcacarccidadecaderacececeredadecacecececes 99
Qy 218	DroserAspleuTyrLeuProProProAspHisGlyAlaProAlaArgGlySerProHis 21
op qu	93 CCCAGTGACCTCTACCTCCCGCCCCCGGACCATGGTGCCCCGGGCCCGTGGCTCCCCCAC 934
2y 22	00 SerGluGlyGlyDysArgSerProGluProAsnLysThrSerValLeuGlyGlyGlyGlu 22
6 q ₀	33 AGCGAAGGGGGCAAGAGTCTCCAGAGCCAAACAAGACGTCGGTCTTGGGTGGTGGTGAGGAG 8/4
∾.	20 AspGly1leGluProValSerProProGluGlyMetThrGluProGlyHisSerArgSer 2239
	73 GACGGTATTGAACCTGTGTCCCCACCCGGGGGCTTGACGGGGGGGG
δy 22	40 AlavalTyrProLeuLeuTyrArgAspGlyGluGlnThrGluProSefArgMeTGlySet 2233
8 qo	/54
2y 22	60 LysSerProGlyAsnThrSerGlnProProAlaPhePheSerLysLeuThrGluSerAsn 2279
	CCC
Oy 22 Db 6	80 SeralametValLysSerLysLysGinGluIleAsnLysLysLeuAsnThrHisAsnArg 2299
2y . 23	00 AsnGluProGluTyrAsnIleSerGlnProGlyThrGluIlePheAsnMetProAlaIle 2319

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                                                                                                      ---AGGACTAAAACATCAAACCGTTACAGCCCAGAATCCCAGGCTCAGTCTGTCCATCAT
                                                                                                                                                                                                                  CAAAGACCÁGGTTCAÁGGGTCTCTCCÁGAAATCTTGTGGACAAATCCÁGGGGÁAGTAGG
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                                                                                                                                                            ----SerGluGlyGlyLys
                                                                                                                                                                                                                                                                        2205 ArgSerProGluProAsnLysThrSerValLeuGlyGlyGlyGluAspGlyIleGluPro
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                                                     LeuArgArgProProSerAspLeuTyrLeuProProAsp-
                                                                                                                                                            2191 GlyAlaPro---AlaArgGlySerProHis-----
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antiinflammatory; cerebroprotective; antilipaemic; antidiabetic; immunouppressive; neuroprotective; notropic; neuroleptic; tranquilizer; osteopathic; antiarthritic; antirheumatic; cytostatic; hepatotropic; virucide; haemostatic; anti-HIV; antihyroid; thyromimetic; anticorpic; chemaclogical; antibacterial; fungicide; antiparasitic; anticonvulsant; thrombolytic; anticoaqulant; anorectic; vasotropic; antiucer; gene therapy; protein replacement therapy; human; gene; ds.
                                                                                                                                           diagnostic and therapeutic polynucleotide; dithp; antiarteriosclerotic;
                                                                                                                        Human diagnostic and therapeutic polynucleotide (dithp), SEQ ID No 61.
2512 ThrLeuSerAspSerGlu 2517
                  7083 ACCCTGTCGGATAGTGAT 7100
                                                           BP
                                                                                                                                                                                                                                                                                                                      16-JAN-2002; 2002US-0349384P.
17-JAN-2002; 2002US-0349413P.
17-JAN-2002; 2002US-0349946P.
                                                           ADE31306 standard; DNA; 2336
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                                                                                                   29-JAN-2004
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田; Gerstin Chinn J, Dufour GE, Jackson JL, by SR, Dam TC, Liu TF, Gerstin Marwaha R, Lan RY, Urashka ME; son O, Yap PE, Amshey SR, Dam Lewis SA, Chen A, Marwaha R, Kolluru V, Panesar IS; Dahl CR, Glect A Yap PE, 2003-636732/60. Tuason O, P-PSDB; ADE31117 Kristnam SR, Peralta CH, ž

Jackson JL;

Gietzen D,

Jones AL,

New human diagnostic and therapeutic polynucleotides and polypeptides, useful for diagnosing, treating or preventing e.g. leukemia, brain cancer, atherosclerosis, AIDS, thyroiditis, infections, obesity, stroke or Alzheimer's.

Claim 1; SEQ ID NO 61; 634pp; English.

The invention relates to a novel isolated human diagnostic and therapeutic polymucleotide (designated dithp). The novel dithp polymucleotide comprises: any of 188 DNA sequences consisting of 195-7798 comprising a naturally defined in the specification; a polymucleotide complementary to comprising a naturally occurring polymucleotide sequence at least 90% identical to the dithp polymucleotide; a polymucleotide complementary to the dithp polymucleotide of the polymucleotide of the dithp polymucleotide of any of the polymucleotides mentioned showe. The dithp polymucleotides have the following activities: antidabetic, antidabetic, antidamentary, cerebroprotective, antilipaemic, antidabetic, immunosuppressive, neuroprotective, notropic, neuroleptic, tranquilizer, osteopethic, antiarthritic, antithheumatic, cytostatic, hepatotropic, virucide, haemostatic, antithromid, thyromimetic dermatological, antibacterial, fungicide, antiparasitic, anticonvulsant, thrombolytic, anticonvulsant, anorectic, vasotropic, and antiuleer. The novel DITHP polymucleotides polypeptide can be used in gene therapy and protein replacement therapy. The dithp polymucleotides polypeptide can be used in gene therapy and protein replacement therapy. The dithp polymucleotides or DITHP protein replacement services associated with the expression of human molecules. In particular, these diseases include cancers (e.g. adenocarcinoma, leukaemia, meaning

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2172 GCAGAGCAGTGCARGATGCARGAGCACAGTGCTGAAAGGCACTTCTTTCTCTAT 2231 229 CCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCCC	Qy 1167 ValLysGlnGluGlnLeuSerProArgGlyGlnAlaGly 		Qy 1207 SerileThrLysGlylleProSerThrArgValProSe:	Qy 1227 GlySerIleThrHisGlyThrProAla Db 3282 GGCTCTATCACTCAGGGCACCCCGGCTCTGCCCCAGAC	Qy 1239 TyrLysGlyThrileThrArgilelleGlyGluAspSe	1259 A 3396 A	Qy 1279 SerTyrGluGlyGlyMetSerValThrGlnCysSerLy:	Oy 1299 GlyProProHisGluThrAlaAlaProLysArgThrTy	Qy 1319 GlyArgalaileSerSerAl. ::	Qy 1333 ArgalalleProProGluArgHisSerProHisHi 	Qy 1352 ArgGlySerIleThrGlnGlyIleProArgSerTyrVa	Qy 1372 ArgArgGluAlaLysLeuLeuLysArgGluGlyThrPrint :::	Qy 1392 AspLeuThrGluAlaTyrLysThrGlnAlaLeuGlyPro	Oy 1412 GluGlyLeuValhalaThrValLysGluAlaGlyArgSe:	Qy 1432 GluLeuArgHisThrProGluLei ::::::	Qy 1447 LysGluGlySerIleThrGlnGlyThrProLeuLysTy. IIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIII	Oy 1467 GlySerLysLysHisAspValArgSerLeuIleGlySe:
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yraspMetMetGluGlyArgVal 1318 ||::: ::|||||| |Argaarcagrgaaggaarara 3557 ::||| rtgacaacaac---tcaggtcaa 3983 erProSerArgLeuAspArgGly 1258 ysGluAspGlyArgSerSerSer 1298 --CGAGAAGGACTAGGAGTCCA 3500 laSerIleGluGlyLeuMetGly 1332 || :::|||||||||::: |caccGrraGaGGGCrGararGc 3617 alGlualaGlnGluaspTyrLeu 1371 |||:::|||||| |CTGAAAGCTTTGAAGATGGCCTT 3731 euProLeuAlaProArgProLeu 1446 :: ::: ||||||::: TGGTCCAGAGCACACGGCCGATA 3926 yraspThrGlyAlaserThrThr 1466 erproGlyArgThrPheProPro 1486 laLeuGluArgAlaCysTyrGlu 1505 :::|||||| rTGGTAGAACGGGGAAATATGAG 4103 .laLeuGlySerValProGlyGly 1206 ::: ||| CTGGCATACCAACAGAGGCTTTG 3341 luglylyslysglyhisvalleu 1278 iisLeuLysGluGlnHisHisIle 1351
||||||||||||| roProProProProSerArg 1391 roLeuLygLeuLygProAlaHis 1411 --ACCAAAGGAAAACCA---TAT 3809 GGCCTAGCAAACTATCCCGTGGA 4043 lyProProGluSerLeuGlyVal 1186 cacaaccreaegercrerredre 3167 erAspserAlalleThrTyrArg 1226 :::||| TGGAGAGCATTCCATCCCTACGG 3281 ----- 1238 ACCTCAAAGAAGGACTGTATTG 3671

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The present invention relates to novel nucleotide sequences, which are differentially expressed in models of osteogenesis upon being put in contact with a stimulator of osteogenesis. The present sequence is one such sequence. This sequence can be used for diagnosing osteoprosis/bone disease in a parient, promoting osteogenesis and/or preventing osteoprosis/bone disease. The present sequence encodes a transcription
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5285 ACGGGAAAGGGAGAAGCGCGAACGTGAGCGCGAGCGTGA 1819 rThrValGluHisAlaPro1leTrpArgProGl ::: 5345 AAGGATCGCTGCTCCCGCTGACCTCTACCTACGAGCAGG 1836 ySerSerGlySerSerGlyGlyGlyGlyGlyGlyGlyGlyGrSerSerSerAr					rLeumetGlubrovalb ACCCCAGATGGATGITT	5717 AAATTTGAGAAGCAGGTCAGCAGCTTAGTGAACAGCAGCAGCAGCAGCAGCAGCAGCAGCAGCAGCAGC	1958 aPheLeuAlaLysProProAlaArgSerGlyLeuGluProAl :::	1978 ySerdluProArgProLeuVal	1987 ovalSerGlyHisAlaThrIleAlaArgThrProAlaLysAs :: see7 aaarCcagaraTGAGGAAGAGCTAAGGACCCGAGGAAGAC	2005 sHisAlaSerProAspProProAlaProProAlaSerAlaSe::::::::::::::::::::::::::::::::::::	2025 sThrGlnSerLysProPheSerIleGlnGluLeuGluLeuAr -:: CTCTCAAAGTTCAGACTCTTCTAG	2045 ySerSerTyrSerProGluGlyValGluProValSerProVa :::::: :::	2065 rHisaspLysGlyLeuProLysHisLeuGluGluLeuAspLy 	2085 uLeuargProLysGlnProGlyProValLysLeuGlyGlyGl 	2105 sLeuArg	2120 uGlnThrAlaProGlyValLysGlyHisGlnArgVa :: CAGTCAGAGGAATGGGACAGGTGCCCAGGACCCATCGACT
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	aArgalaLeuGluArgAlaCysTyrGluGluSerLeuLysSerAr	4484 GCACACGTGGTGAGCTCTGGCCCCTCCGTTCTCAGGTCTACACTTCA 4534 1531 oGluLeuGlyLysProArgGlnSerProLeuThrTyrGluAspHisGlyAla 1548	1549PropheAlaGlyHisLeuProArgGlySerProValThrMetArgGluProTh 1566 :::	1566 rProArgLeuGlnGluGlySerLeuSerSerLysAlaSerGlnAspArgLysLe 1585	1585 uThrSerThrProArgGluIlealaLysSerProHisSerThrValPr 1601 	1601 oGluHisHisProHisProIleSerProTyrGluHisLeuLeuArgGlyValSerGlyVa 1621 	1AspLeuTyrArgSerHisT 		rProHisLeuTyrProProTyrLeuIleArgGlyTyrProAspThrAlaAlaLeuGluAs 	pTyrlleThrSerClnGlnMetHisHisAsnThrAl	aThralaMetalaGlnargalaAspWetLeuArgGlyLeuSerProArgGluSerSerLe	ualaLeuAsnIyrAlaalaGlyProArgGlyIleIleAspLeuSerGlnValProHisLe	5060 GGGGCTCCCTTACCCAGCTACAAGAGGAATCATTGACCTGACC	rbeuproThrAlaProGlnProPheSerSerArgHisSerSerBerProLeuSerPr	1779 oGlydlyProThrHisLeuThrLysProThrThrThrSerSerGluArgGluArgAs 1799	
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GlyThrGluGlnSerSerGl 1836 ||||:::||||||| |GGTTCAGAACAG----- 5396 AlaserSerProserLysGl 1978
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TCAGCCCTTCCAAGTGGCAA 5836 ArgSerLeuGlyTyrHisGl 2045 ValSerSerProSerLeuTh 2065 LysSerHisLeuGluGlyGl 2085 ArgProAlaSerHisSerHi 1856 AlavalGluProSerLysPr 1895 ||||||: AGTCAAGGCCTGCCAGCCTC 5599 :::||| GTGGATGCTGCAGCTTCTGC 5656 ArgAlaAspThrGlyHisAl 1958 CAGCTAGAGCAGAAAACCT 5776 AAAGATAAAGGGCCTCCTCC 5896 Asn-----LeuAlaProHi 2005 ||| :: ACTACCATTACTGCAGCTAA 5956 SerAspProHisArgGluLy 2025 AAGGATGCGAGGGACGTGG 6016 SCAAATCAAGCAGAAAATGA 6178 3luAlaAlaHisLeuProHi 2105 ProSerSerProLeuLe 2120 Ala---LeuGlnGlnArgPr 1875 ::::::|||::: ------Argproalaal 1911 LeuAspGlyValTyrProTh 1931 1944 SAAGCTGCCAGGTTAGAAGA 5716 ...---- 1987 SAGCGTGAACGTGAACGCGA 5344

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                                                                                                                         The invention describes a method of diagnosing breast cancer in subject comprising contacting biological sample from subject with at least two addifferent breast cancer-associated polypeptides (I) encoded by nucleic acid molecules (II) comprising sequence chosen from 42 fully defined sequences as given in specification, determining specific binding between (I) and agents in sample, where presence of the binding is diagnostic for breast cancer. The method is useful for diagnosing breast cancer in a gubject. The sample is blood, lymph node fluid or breast discharge fluid
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Human; ss; granulocytic cell; DNA chip; bacterial infection; viral infection; parasitic infection; protozoal infection; fungal infection; sterile inflammatory disease; psoriasis; rheumatoid arthritis; glomerulonephritis; asthma; thrombosis; cardiac reperfusion injury; renal reperfusion injury; ARDS; andlt respiratory distress syndrome; inflammatory bowel disease; Crohn's disease; ulcerative colitis; periodontal disease; granulocyte activation; chronic inflammation; allergy.
                                                                                                           Human cDNA differentially expressed in granulocytic cells #876
                                                                                                                                                                                                                                                                                                                             Vockley
                                                                                                                                                                                                                                                                                                                             Yamaga S,
2517
                                                     ABK84305 standard; cDNA; 5989 BP
                                                                                                                                                                                                                                                                                                                           Beazer-Barclay Y, Weissman SM,
                                                                                                                                                                                                                                                                      03-OCT-2001; 2001WO-US030821.
                                                                                                                                                                                                                                                                                        03-OCT-2000; 2000US-0237189P.
           7528 ACACTCTCGGACAGCGAG
2512 ThrLeuSerAspSerGlu
                                                                                           (first entry)
                                                                                                                                                                                                                                                                                                          (GENE-) GENE LOGIC INC
                                                                                                                                                                                                                                                                                                                                             WPI; 2002-435328/46.
                                                                                                                                                                                                                                  WO200228999-A2
                                                                                                                                                                                                                 Homo sapiens
                                                                                         14-AUG-2002
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Detecting granulocyte activation by detecting differential expression of genes associated with granulocyte activation, which serves as diagnostic markers that is useful for monitoring disease states and drug toxicity.

Claim 1; SEQ ID NO 876; 114pp; English.

The invention relates to detecting (M1) granulocyte (GC) activation (GCA), by detecting the level of expression of gene(s) (GS) identified by DNA chip analysis as given in the specification, and comparing the expression level to an expression level in an unactivated GC, where differential expression of GS is indicative of GCA. Also included are condulating (M2) GA by conteating GC with an agent that alters the expression of at least one gene in GS; (2) screening (M3) for an agent capable of modulating GCA or an inflammation (especially chronic) in a capable of modulating GCA or an inflammation (especially chronic) in a crissue, an allergic response in a subject, exposure of a subject to a pathogen or sterile inflammatory disease using the gene expression of the subject, exposure of a subject to a pathogen or sterile inflammatory disease, by detecting the level of expression in a sample of the tissue of gene(s) from GS, where the level of expression of the gene is indicative of inflammation; (4) treating (M5) an inflammation of the gene is indicative of inflammation with an expression of sepecially chronic) or in a tissue, an allergic response in a subject, exposure of a subject to a pathogen or sterile conflammation of sepecially chronic) or in a tissue, an allergic response in a subject, exposure of a subject to a pathogen or sterile conflammation in a tissue, M4 is useful for detecting GCA, M2 is useful for modulating GA, M3 is useful conflammation in a tissue, an allergic response in a subject, conflammation of gene (s) from GS in the tissue, m1 inflammation (especially chronic) in a tissue, an allergic response in a subject, conflammation disease, conflammation disease, conflammation and attitudes and allergic response in a subject, conflammation in a tissue, m4 is useful for detecting an attitude of a subject to a pathogen or sterile inflammation disease (e.g. exposure of a subject to a pathogen or sterile inflammation and an exposure of a subject, renal reperfusion injury, renal reperfusion injury, renal

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                             infection,
                      infection, viral infection, parasitic infection, protozoal infectio fungal infection and MS is useful for treating one of the above conditions. The present sequence represents a gene differentially expressed in granulocytes. Note: The sequence data for this patent not form part of the printed specification, but was obtained in electronic format directly from WIPO at ftp.wipo.int/pub/published_pct_sequences
disease, ulcerative colitis, periodontal disease; also bacterial
                                                                                                                                                                                                                                                                          T; 0 U; 0 Other;
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	360 GlyLeuSerMetSerAlaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeu 379 	380 SerGluGlnGluAsnLeuGluLysGlnMetArgGlnLeuAlaValIleProProMetLeu 399 	400 TyrabplaabglinginargileLysPheileasnMetasnGlyLeuMetalaaspPro 419 	420 MetlysValTyrLysAspArgGlnValMetAsnMetTrpSerGluGluGlnGluLysGluThr 439 	440 PheArgGluLysPheMetGlnHisProLysAanPheGlyLeuIleAlaSerPheLeuGlu 459 	ArglysThrValAlaGluCysValLeuTyrTyrTyrLeuThrLysLysAsnGluAsnTyr	AGAAAGACGSTCGCTGAGTGTGTCCTCTATTACTACCTGACCAGGAAAAAAAA	AAGAGCTTGGTGAGGGGGGGGGTATCGGCGCCGTGGCAAGAGCCAGGCAGCAGCAGCAGCAGCAGCAGCAGCAG					1885 GGCCGTATCACGCGCTCCATGGCCAACGAGGCCATGAGGAGACAGCCACGCCACAGG 1944 600 GlnSerAlaGluLeuAlaSerMetGluLeuAsGGluSerSerArgTrpThrGluGluGlu 619	1945 CAAAGTTCAGAGCTTCCATGAGATGAAAGTTCTCGCTGGACTGAGGAAGAG 2004 620 MetGluThrAlaLysLygGlyLeuLeuGluHigGlyArgAsnTrpSerAlaIlaAlaArg 639	20	640 MetValGlySerLysThrValSerGlnCyeLysAsnPheTyrPheAsnTyrLysLysArg 659	660 GlnAsnLeuAspGluIleLeuGlnGlnHisLysLeuLysMetGluLysGluArgAsnAla 679	2185 CGGAGGAAGAAGAAGAAGCCCCAGCTGCGGCGAGGAGAGAGA

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  New SMRTe proteins and nucleic acids, useful in gene therapy, predictive medicine, therapeutic or prophylactic treatment, chromosome mapping, tissue typing and in forensic biology.
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                                             AlaGlyProHisHisAlaTrpAspGluGluProLysProLeuLeuCysSerGlnTyrGlu
                                                                     GCTGGTCCCCCACCACGCCTGGGATGAGGAGCCCAAGCCACTGCTGTTTCACAGTATGAG
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                                                                                                                                   ACA62451 standard; cDNA; 8544 BP
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ProArgArgArgAlaLysGluSerLysValArgGluTyrTyrGluLysGlnPheProGlu

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Length:
Matches:
Conservative:
Mismatches:
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                                  ss; gene; SMRTe; nuclear receptor corepressor; gene therapy;
typing; cancer.
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Sequence 7386 BP; 1735 A; 2520 C; 2046 G; 1085 T; 0 U; 0 Other;

Alignment Scores:

Qy 2112 rGlnProSerSerSerProLeuLeuGlnThrAlaProGlyValLysGlyHisGlnArgVa	2132 IValThrLeualaglnHisIleSerGluValIleThrGlnAspTyrThrArgHisHi 	2152 oglnGlnLeuSerAlaProLeuProAlaProLeuTyrSerPheProGlyAlaSerCy 	2172 oValleuAspleuArgArgProProSezAspleuTyrLeuProProProAspHisGl	2192 aProAlaArgGlySerProHisSerGluGlyGlyLysArgSerProGluProAe	2212 rSerValLeuGlyGlyGlyGluAspGlyIleGluProValSer	Oy 2232 rGlubroGlyHisSerArgSerAlaVaTYrProLeuLeuTyrArgAspGlyGluclnTh 	2252	2272 eSerLysLeuThrGluSerAsnSerAlaMetValLysSerLysL 	2292 sLysLeudsnThrHisAsnArgasnGluProGluTyrAsnIleSerGlnProGlyTh	2312 ullePheAsnWetProAlaIleThrGlyThrGlyLeuMetThrTyrAr 	2332 IGInGludisAlaSerThrAsnMetGlyLeuGluAlaIleIleArgLysAlaLeuMe 	2352 yLysTyrAspGlnTrpGluGluSerProProLeuSerAlaAanAlaPheAsnProL 	2372 nAlaSerAlaSerLeuProAlaAlaMetProIleThrAlaAlaAspGJyArgSerA 	Qy 2392 sThrLeuThrSerProGlyGlyGlyLysAlaLysValSerGlyArgProSerSerAr 	2412 gLysAlaLysSerProAlaProGlyLeuAlaSerGlyAspArgProProSerV	7422	Oy 2452 pArgProSerSerAlaGLySerThrProPheProTyrAanProLeulleMetArgLeuGL
5322 GGACCTGTCCCAAGTGCCACCTGCCTGTGCTCGTGCCCCGGACACCCAGGCACCCCAGC 5381 1752 aThrAlaMetAspArgLeuAlaTyrLeuProThrAlaProGInProPheSerSerArgHi 1772	SERSERSERPROLEUSERPROGLYGLYPROTHRHISLEUTHRUSPROTHRTHRTHRIBE	r L	1812 sSerIleLeuThrSerThrThrThrValGluHisAlaProIleTrpArgProGlyThrGl 1832 	1832 uGlnSerSerGlySerSerGlyGlyGlyGlyGlyGlyGlyGrSerSerArgProAl 1852 	1852 aSerHisSerHisAlaHisGlnHisSerProlleSerProArgThrGlnAspAlaLeuGl 1872 	1872 nGlnargproSerValLeuHisasnThrGlyMetLysGlyIleIleThralaValGluPr 1892 		1912 rPheProProAlaThrHisCysProLeuGlyGlyThrLeuAspGlyValTyrProThrLe 1932 	1932 UMetGluProValLeuLeuProLysGluAlaProArgValAlaArgProGluArgProAr 1952 	1952 gAlaAspThrGlyHisAlaPheLeuAlaLy8ProProAlaArgSerGlyLeuGluProAl 1972 	1972 aSerSerProSerLysGlySerGlubroArgProLeuValProProValSerGlyHisAl 1992 	1992 aThrileAlaArgThrProAlaLysAsnLeuAlaProHisHisAlaSerProAspProPr 2012 	2012 oAlaProProAlaSerAlaSerAspProHisArgGluLysThrGlnSerLysProPheSe 2032		yvalgluprovalSerProvalSerSerProSerLeuThrHisAspLysGlyLeuProLy	2072 SHisLeuGluGluLeuAspLysSerHisLeuGluGlyGluLeuArgProLysGlnProGl 2092 	2092 yProValLysLeuGlyGlyGlualaalaHisLeuProHisLeuArgProLeuProGluSe 2112

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997 ATCTTGTACTTCAAGAGGAATCACGCTCGGAAACAATGGGAGCAGAAGTTCTGCCAG 1056 OY 301 AKGTYKASDGInLeuMetGluAlaLeuGluLysLysValGluArgIleGluAsnAsnAsnPro 320	ArgArgArgAlaLysGluSerLysValArgGluTyrTyrGluLysGlnPheProGluIle 3	rrGly 360 rrGGG 1233	361 LeuSerMetSerAlaAlaArgSerGluHisGluValSerGluIleIleAspGlyLeuSer 380 	281 GluGlnGluAsnLeuGluLysGlnMetArgGlnLeuAlaValIleProProMetLeuTyr 400	401 AspalaaspGlnGlnargIleLysPheIleAsnMetAsnGlyLeuMetAlaAspProMet 420	rPhe 440 CTTC 1473	ArgGluLysPheMetGlnHisProLysAsnPheGlyLeuIleAlasSerPheLeuGluArg 4 	LysThrValAlaGluCysValLeuTyrTyrTyrLeuThrLysLysAsnGluAsnTyrLys 4		uLys 5	ASDGLULYSGLULYSGLULYSGLULASGLUCHISCCCCGCAGCAGCAGGAGGAGGAGGAGGAGGAGGAGGAGGA	ו נט רו	BG1y 580 AGGC 1893	ArgileThrArgSerMetAlaAsnGluAlaAsnSerGluGluAlaileThrProGlnGln 6	SeralaGluLeualaSerMetGluLeuasnGluSerSerArgTrpThrGluGluGluMet 62	AGCGCCGAGGCIGGCCICCCAIGGAGGCIGAAIGAGAGGIICICGCIGGACAGAAAAI GluIhralaLyBLyBClyLeuLeuGluHiBGlyArgABnTrpSerAlaIleAlaArgMe 	, d – t,

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                                             7266 AGTGCACTCGGAGGGAGACTGCAACCGCCGGACGCCCCTCACCACCGCGTGTGGGAGGA
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tissue typing; cancer.
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acid expression or activity, such as cancer. The present sequence represents cDNA encoding the human nuclear receptor corepressor SMRTe
                                                      Sequence 8686 BP; 1881 A; 3069 C; 2562 G; 1174 T; 0 U; 0 Other;
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Conservative:
Mismatches:
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                                                                                                                                                                                                                                                                                                                                                                                                                                       GluGluProLysProLeuLeuCysSerGlnTyrGluThrLeuSerAspSerGlu 2517
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                                                                                                                                                                                    nucleic
amplifying of SMRTe nucleic acid molecules. The polypeptides are useful as immunogens to raise anti-SMRTe antibodies. The SMRTe molecules are useful as targets for discovering and developing modulating agents to regulate a variety of cellular processes, in screening assays, in predictive medicine, in therapeutic or prophylactic treatment, in chromosome mapping, tissue typing and in forensic identification of a biological sample. Modulators of SMRTe are useful for treating or preventing a condition associated with aberrant SMRTe protein or nuclead expression or activity, such as cancer. The present sequence
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     MetSerGlySerThrGlnLeuValAlaGlnThrTrpArgAlaThrGluProArgTyrPro
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Matches:
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2020 AspPrOHisArgGluLysThrGlnSerLysProPheSerIleGlnGluLeuGluLeuArg 2039 [6361 CTCCAGCCGCCCCAGGGGTCACCAGCGGGTCCCCCCCTGGCCCAGCACATC 6420 2140 SerGluVallleThCGllnAspTyrThrArgHisHisProGlnGlnLeuSerAlaProLeu 2159 [11] [11] [11] [11] [11] [11] [11] [11]	SerGluGlyGlyLyshrgSerProGluProAsnLysThrSerValLeuGlyGlyGlyGlu	6781 AAGTCTCCAGGCAACCAGCCAGCCAGCCTTCTTCAGCAAGCTGACCGAGAGCAAC 6840 2280 SerAlaMetValLysSerLysLysGlnGlulleAsnLysLysLeuAsnThrHisAsnArg 2299 2280 SerAlaMetValLysSerLysLysGlnGlulleAsnLysLysLeuAsnThrHisAsnArg 2299 [6961 ACCGGACACGACCTATAGACCTATAGACCCGAGCGCGACAGGACATGCCAGCACCACA 7020 2340 MetGlyLeuGlualatielleacypsalaleuMetGlyLysTyraspGlnTrpGluGlu 2359 2340 MetGlyLeuGlualatielleacypsalalacumetGlyLysTyraspGlnTrpGluGlu 2359 2361 ATGGGCTGGAGGCCATAATAGAAAGGCACTCATGGGTAATATGAAAGAGCACTCATGGGTAATATGAAGAGG 7080 2360 SerProProLeuSerAlaAshalaPheAsnProLeuAsnAlaSerAlaSerLeuProAla 2379 2360 SIPHIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIIII
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ThrTyrProHisLeuTyrProProTyrLeuIleArgGlyTyrProAspThrAlaAlaLeu 167 ACCTACCCGCACACCTACCCCTACTCTCGCGGCTACCCCGACACGCGGCTG 504 GluAsnArgGlnThrIleIleAsnAspTyrIleThrSerGlnGlnMetHisHisAsnThr 169 GluAsnArgGlnThrIleIleAsnAspTyrIleThrSerGlnGlnMetHisHisAsnThr 169 AlaThrAlaMetAlaGlnArgAlaAspMetLeuArgGlyLeuSerProArgGluSerSer 171 GCACCGCCATGGCCCATCATCATCACATCACTCGCACGCGAGATGCACACACA		Hisdin 185; Hisdin 185; CACCAG 558 CACCAG 558 CHUHIS 187; CTTCAC 564; CTTCAC 564; CTTCAC 564; CTTCAG 570;		1900 Lyshaniaystrontonianiastis (http://linkinianiastis.com/pourseling) 5881 CTCGCCAAGCCCCGCTCCGGGCTGGAGCCCGCCTCCTCCCCCCAGCAAGGCTCG 5940 1980 GlubroArgProLeuValProProValSerGlyHisAlaThrileAlaArgThrProAla 1999 1981 GAGCCCCGGCCCCTAGTGCTCTCTGGCCACCCTCCCCCCGCACCCCTGCG 6000 2000 LyshanieuAlaProHisHisAlaSerProAspProProAlaProProAlaSerAlaSer 2019

130 Proprofit GluthAlabiaProLyshaght-YirlapkethetolidyArgyaldy 1119
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Acc78130 Nucleotid	Aah23705 Human tum	Abs56090 S. venezu	Aaz87318 S. venezu	Aaz87285 S. venezu	Aaa75633 Nucleotid	Aaz56001 Recombina	Ada09418 Cosmid pK	Continuation (7 of	Aaq46806 eryA regi	Abv75361 Human KRC	Aah23695 Human tum	Acd13385 Human DNA	Acc50291 Breast ca	Ach32484 Human end	Continuation (19 o	Aac76700 Human ORF	Continuation (19 o	Abq61152 MUC5B par
ACC78130	AAH23705	ABS56090	AAZ87318	AAZ87285	AAA75633	AAZ56001	ADA09418	AA199683 06	AAQ46806	ABV75361	AAH23695	ACD13385	ACC50291	ACH32484	AAI99683 18	AAC76700	AA199682 18	ABQ61152
34094 8	906 4	38506 7	778 3	37948 3	38506 3	506 3	38506 7	10000 4	29879 2	8546 7	115 4	11145 7	227 7	199 8	10000 4	11680 3	10000 4	13715 6
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ALIGNMENTS

AAC74783 standard; cDNA; 8564 BP (first entry) 08-FEB-2001 AAC74783;

Human ORFX ORF338 polynucleotide sequence SEQ ID NO:675.

Human; open reading frame, ORFX; detection; cytostatic; hepatotropic;
wulnerary; antipsoriatic; antiparkinsonian; nootropic; neuroprotective;
wulnerary; antipsoriatic; antiparkinsonian; nootropic; neuroprotective;
anticonvulsant; osteopathic; antiarthritic; immunosuppressant; cardiant;
immunostimulant; thrombolytic; coagulant; vasotropic; antidiabetic;
Mypotensive; dermatological; immunosuppressive; antiinflammatory;
antiviral; antibacterial; antifungal; antitheumatic; antithyroid;
antianaemic; gene therapy; cancer; proliferative disorder; hypertension;
metrodegenerative disorder; osteoarthritis; graft vs host disease;
heurodegenerative disorder; osteoarthritis; hypothyroidism; SCID; ADDS;
cholesterol ester storage; systemic lupus erythematosus; infection;
severe combined immunodeficiency; malaria; autoimmune disorder; asthma;
allergy; applastic anaemia; nocturnal haemoglobinuria; burn; wound;
bone damage; cartilage damage; antiinflammatory disease; coagulation;
whenombosis; contraceptive; ss. RESULT 1
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WO200058473-A2.

05-OCT-2000

31-MAR-2000; 2000WO-US008621.

02-APR-1999; 99US-0127636P. 05-APR-1999; 99US-0127728P. 30-MAR-2000; 2000US-00540763. 99US-0127607P. 31-MAR-1999;

(CURA-) CURAGEN CORP.

Shimkets RA, Leach M;

WPI; 2000-602362/57. P-PSDB; AAB40574.

Novel nucleic acids and peptides derived from open reading frame X, useful for treating e.g. cancers, proliferative disorders, neurodegenerative disorders and cardiovascular disease.

200

181 ArgGluIleThrMetValGluGlnGlnIleSerLysLeuLysLysLysGlnGlnGlnLeu 541 CGAGAGATCACCATGGTAGAGCAGCAGATCTCTAAGCTGAAGAAGAAGCAGCAACAGCTG

900

Claim 5; Page 779-784; 5507pp; English

AAC74446 to AAC77606 encode the proteins given in AAB40237 to AAB43397,
which represent the human ORFX open reading frames 1 to 3161. The ORFX
completes have activities such as: cytostatic; hepatotropic; vulnerary;
antipsoriatic; antiparkinsonian; nootropic; neuroprotective; osteopathic;
anticonvulsant; antiarthriti; immunosuppressant; immunostimulant;
cardiant; thrombolytic; coagulant; vasotropic; antidiabetic; hypotensive;
dermatological; immunosuppressive; antidiabetic; hypotensive;
dermatological; antirheumatic; antithyroid; and antianaemic.
cardiant; thrombolytic; coagulant; vasotropic; antidiabetic; hypotensive;
dermatological; antirheumatic; antithyroid; and antianaemic.
cardiant; thrombolytic; coagulant; vasotropic; antidiabetic; hypotensive;
cardiant; thrembolytic; coagulant; antithyroid; and antianaemic. The
sequences can be used for determining the presence of or predisposition.
CCC o, or preventing or treating pathological conditions associated with an
CORFX-associated disorder. The nucleic acids can be used to express ORFX
proteins in gene therapy vectors. The proteins and nucleic acids may be
disorders osteoarthritis, graft vs host disease, cardiovascular disease,
cdiabetes mellitus, hypertension, hypothyroidism, cholesterol ester
storage, systemic lupus erythematosus, severe combined immunodeficiency
CC (SCID), AIDS, viral, bacterial or fungal infection, malaria, autoimmune
cdisorders, asthma, allergies, aplastic anaemia, burns, wounds, bone and
cartilage damage, nocturnal haemoglobinuria, antiinflammatory disease; to
enhance coagulation; to inhibit thrombosis; and as a contraceptive

Seguence 8564 BP; 1859 A; 3034 C; 2532 G; 1139 T; 0 U; 0 Other;

	8564	2512		4		.
	Length:	Matches:	Conservative:	Mismatches:	Indels:	Gaps:
	0	13178.50	808.66	99.768	99.728	ĸ
Alignment Scores:	Pred. No.:	Score:	Percent Similarity:	Best Local Similarity:	Query Match:	DB:

US-09-522-753-5 (1-2517) x AAC74783 (1-8564)

셤 ò 용 à 셤 ઠે 유 ò g ò 셤 ò 셤 ò . සු ò 셤 ò 셤

н	MetSerGlySerThrGlnLeuValAlaGlnThrTrpArgAlaThrGluProArgTyrPro 20
П	ATGTCGGGCTCCACACACATGTGGCACAGACGTGGAGGCCCACTGAGCCCCGCTACCCG 60
21	ProHisSerLeuSerTyrProValGln1leAlaArgThrHisThrAspValGlyLeuLeu 40
61	CCCCACAGGCTTTCCTACCCAGTGCAGATCGCCCGGACGCACACGGACGTCGGGCTCCTG 120
41	GluTyrGlnHisHisSerArgAspTyrAlaSerHisLeuSerProGlySerIleIleGln 60
121	GAGTACCAGCACCACTCCCGCGACTATGCCTCCCACCTGTCGCCGGGCTCCATCATCACAC 180
61	ProGlnArgArgAroSerLeuLeuSerGluPheGlnProGlyAsnGluArgSerGln 80
181	CCCCAGGGGGGGGGCCCTCCCTGCTGTCTGAGTTCCAGCCGGGAATGAACGGTCCCAG 240
81	GluLeuHisLeuArgProGluSerHisSerTyrLeuProGluLeuGlyLysSerGluMet 100
241	GAGCTCCACCTGCGGCCCAGAGTCCCCACTCATACCTGCCCCGAGCTGGGGGAAGTCAGAGATG 300
101	GluPheIleGluSerLysArgProArgLeuGluLeuLeuProAspProLeuLeuArgPro
301	GAGTTCATTGAAAGCAAGCGCCCTCGGCTAGAGCTGCTGCTGCCTGC
121	SerProLeuLeualaThrGlyGlnProAlaGlySerGluAspLeuThrLysAspArgSer 140
361	TCACCCCTGCTGGCCACGGGCCAGCCTGCGGGATCTGAAGACCTCACCAAGGACCGTAGC 420
141	LeuThrGlyLysLeuGluProValSerProProSerProProHisThrAspProGluLeu 160
421	CHINITION CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROLL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL CONTROL
161	GluLeuValProProArgLeuSerLysGluGluLeuIleGlnAsnMetAspArgValAsp 180
481	GAGCTGGTGCCCCCCCCCCCCCCCCCCCCCCCCCCCCCC

Ab234833 Coding se Aaa60629 Human HNR Adc31306 Human dia Aca57401 Human adi Ab103131 Drosophil	Aaa02670 Human col Aca57524 Human adi Ade76358 Human BSK Aai97539 Human neu Aca57537 Human adi Ab44934 CNNA enco	NA sequorangin ctinoid crangin orangin	Add5815 Micromono Add5810 Micromono Aaf67220 Novel hum Continuation (9 of Aah79277 Streptomy Continuation (9 of Aaa83201 S. avermi Aat80414 Platenoli	E 600000	Aat80413 Tylactone Continuation (40 o Aad54645 Streptomy Adc26813 Sorangium Aat80415 Hybrid sr Continuation (38 o Aad17184 Streptomy Aad17184 Streptomy Continuation (40 o Ach20441 Human adu Aat93095 Streptomy Aat93095 Streptomy Aav25925 Streptomy	Aat 66769 Sorangium Aat 89956 Sorangium Aaa 75299 DNA seque Aav 6287 A secue Aaa 92302 S. avermi Aah 79278 Streptomy Continuation (3 of Aaf 88313 S. spinos Aaf 88315 S. spinos Aaf 88315 S. spinos Aat 88316 S. spinos Aat 88316 S. spinos Acontinuation (4 of	Aa25887 Sorangium Aaa59349 Sorangium Aad5817 Micromono Aa161224 Actinosyn Aaf30757 Micromono Continuation (13 o Aaq73500 DNA Aaf88338 S. spinos Aaf88315 Streptomy	AA28/1285 venezu Abx04971 S. cinnam Continuation (7 of Continuation (7 of Continuation (13 o Aa161224 Actinosyn Aaf88339 S. spinos Aaz30163 Complete
		AAS17357 AAS17367 ADC26979 ADC26995 9 ADC26991			2 AATB0413 4 AAI99683_39 AAD546455 2 AATB0415 2 AATB0415 3 AAA09469 4 AAD17186 4 AAD17186 4 AAD17186 2 AAT93095 2 AAV2925			AAZ8728 ABX0497 AAI9968 AAI9968 AAL6122 AAF8833 AAF8833
4 7940 6 7900 2 2336 4 650 2 10910	872 3 555 7 718 4 710	33.529 33.529 44 1922 46 7251	3 13416 3 60196 2 427 2 110000 2 30690 1 44347	1 44377 1 75236 1 110000 0 58857 0 110000 0 110000	0 43280 0 113000 0 11358 0 11358 0 13887 0 10000 0 55140 0 125401 0 125401 0 125401 0 125401 0 125401 0 125401 0 24379	.9 28598 .9 28958 .9 49377 .8 31422 .8 110000 .8 110000 .8 50000 .8 50000	7 68750 7 71989 7 11238 7 82746 7 147981 7 14000 7 14775 7 15872	15872 103599 110000 110000 182746 16767 34094
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GenCore version 5.1. Copyright (c) 1993 - 2004 Comp	nucleic search, using frame_plus_p2n April 15, 2004, 22:22:55 ; (withou	US-09-522-753-5 13215 1 MSGSTQLVAQTWRATEPRYP		of hits satisfying chosen parameters of length: 0 eq length: 2000000000 eq length: 2000000000 eq length: 2000000000 eq length: 2000000000 eq length: 2000000000 eq length: 2000000000000000000000000000000000000	10,40,114,040	N. Geneseq_29Jan04:* 1. geneseqn190s:* 2. geneseqn1990s:* 3. geneseqn2000s:* 4. geneseqn2001as:* 5. geneseqn201bs:* 6. geneseqn2001bs:* 7. geneseqn2003as:* 9. geneseqn2003as:* 9. geneseqn2003as:*	is the number of results predicater than or equal to the score erived by analysis of the total SUMMARIES Query Match Length DB ID	5 99.7 8564 3 AAC74783 8 98.2 7521 7 ACA62250 8 98.2 8686 7 ACA62459 5 82.0 7386 7 ACA62451 5 82.0 8544 7 ACA62451 15 59.7 5989 6 ABK84305 7.3 31.7 7780 3 AAA60630
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